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#### (57) Abstract

The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of Chlamydia trachomatis, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences, as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the Chlamydia trachomatis genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing Chlamydia trachomatis infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular Chlamydia trachomatis, infections.

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# CHLAMYDIA TRACHOMATIS GENOMIC SEQUENCE AND POLYPEPTIDES, FRAGMENTS THEREOF AND USES THEREOF, IN PARTICULAR FOR THE DIAGNOSIS, PREVENTION AND TREATMENT OF INFECTION

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The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of *Chlamydia trachomatis*, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the *Chlamydia trachomatis* genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing *Chlamydia trachomatis* infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular *Chlamydia trachomatis*, infections.

The genus Chlamydia is composed of four species: Chlamydia psittaci, Chlamydia 20 pecorum, Chlamydia pneumoniae and Chlamydia trachomatis.

Chlamydia psittaci comprises numerous species, whose hosts are terrestrial vertebrate animals as well as birds and occasionally humans;

Chlamydia pecorum is a pathogen of ruminants;

Chlamydia pneumoniae is responsible for pneumopathies, for sinusitis and for arterial impairments in humans;

Chlamydia trachomatis (Ct) is responsible for a large number of human diseases:

- eye diseases: conventional trachoma, nonendemic trachoma, paratrachoma, inclusion conjunctivitis in neonates and in adults;
- genital diseases: nongonococcal uretritis, epididymitis, cervicitis, salpingitis, perihepatitis and bartholinitis as well as pneumopathy in breast-feeding infants;
  - systemic diseases: venereal lymphogranulomatosis (LGV).

These diseases affect a very large number of women and men [more than 600 million individuals are trachoma carriers and there are more than 90 million cases of genital *Chlamydia* infections] worldwide. Accordingly, basic and applied research which makes it possible to understand the physiopathology linked to this bacterium is very important for public health. (Raulston JE., 1995; Hackstadt T. et al., 1996).

Eye impairments due to Chlamydia trachomatis cause trachoma and inclusion

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conjunctivitis. Trachoma is a chronic conjunctivitis. It is the major cause of curable eye diseases leading to blindness. It is estimated that 20 million cases of loss of sight are due to it worldwide. Moreover, inclusion conjunctivitis is an eye inflammation which is caused by *Chlamydia trachomatis* and is transmitted by the venereal route. Inclusion conjunctivitis affects adults and neonates exposed to genital secretions.

Two types of eye disease caused by agents of the species Chlamydia trachomatis can be distinguished. The conventional trachomatous disease is found in endemic regions; transmission occurs from eye to eye and through the hands, or it can be passed on by flies. In nonendemic regions, transmission occurs through the genital apparatus; it usually only causes conjunctivitis, most often 10 without associated keratitis; it is rare for a pannus or for scars similar to those in trachoma to develop. This conjunctival impairment is called paratrachoma to differentiate it from the conventional endemic trachoma which is transmitted by the ocular route. The seriousness and the number of cases of trachoma have decreased over the last forty years. This is related to the improvement in hygiene and living conditions. However, trachoma remains the principal cause of avoidable blindness in Africa, in the Middle East and in some regions of Asia. The transmission of the endemic disease occurs in particular through close personal contact, in regions where a secondary exposure exists in a repeated form. Often, the infection is also latent. In some industrialized countries, such as the United States, a mild form of trachoma still exists in some ethnic groups. Sometimes, a tardive trachoma may be found following an immunosuppressive treatment. The eye impairments caused by Chlamydia trachomatis, such as inclusion conjunctivitis and paratrachoma, are also a complication due to a common venereal infection. These infections are not very frequent; they occur most often in young adults. The eye impairments in neonates are produced during the passage through the maternal genital routes during childbirth. Theoretically, endemic trachoma and inclusion conjunctivitis in adults appear in the form of conjunctivitis, the latter being characterized by the presence of lymphoid follicles. In regions where the endemic disease is serious, the disease often starts before the age of 2 years and reinfection is frequent. Superficial neovascularization is added, in this case, to leukocytic infiltration. The conjunctival scars will then cause trichiasis and entropion. The eroded comea will become a carrier of a corneal ulcer of bacterial origin. The scar on the cornea causes blindness. Impairment of the lachrymal glands gives a picture of dryness of the comea. Xerosis becomes complicated with secondary bacterial ulcer. In regions where trachoma is endemic, the infectious process disappears towards the age of fifteen. The scars then progress to blindness, which affects almost exclusively adults. In regions where exposure is lower, the infectious process is, in this case, less rapid and adults are carriers of a chronic disease.

Positive diagnosis of trachoma can be most often established by clinical observation:

lymphoid follicles are visible on the upper tarsal conjuctiva; conjunctival scar is typical. Vascular

pannus exists. In endemic regions, clinical diagnosis is often sufficient. However, isolated cases of inclusion conjunctivitis must be the subject of a differential diagnosis, in particular to distinguish viral conjunctivitis.

Public health measures against the endemic form of the disease provide for mass treatments with tetracycline or erythromycin collyria of all children. The treatment may also provide for surgical correction of the lesions. The other conjunctival impairments respond well to general treatments with tetracyclines or erythromycin. The prevention of trachomatous disease by health measures and by improving living standards is sufficient. Furthermore, to avoid the spread of trachoma, antibiotic collyria may be used.

The role of *Chlamydia trachomatis* in a number of genital impairments has been demonstrated over the last three decades. *Chlamydia trachomatis* is responsible in this case for a pathology which may be superposed on the impairments observed with *Neisseria gonorrhoeae*. The pathologies for which *Chlamydia trachomatis* may be responsible at the genital level are acquired by the venereal route and are a major source of sexually transmitted diseases.

The epidemiology of Chlamydia trachomatis genital infections shows each year more than 4 million new cases in the United States, and more than 3 million new cases in Europe. Like the other venereal infections, Chlamydia trachomatis affects young subjects. There is a direct relationship between the number of sexual partners and the frequency of the disease. For example, the frequency of Chlamydia trachomatis appears to be five to ten times higher than that of Neisseria gonorrhoeae in pregnant women. The Chlamydia trachomatis infection is probably more discreet than its Neisseria gonorrhoeae homologue. This relative clinical silence, estimated in women at 50% or even 70% of infections, explains why the total morbidity of Chlamydia trachomatis conditions is high. Diagnosis must therefore be requested in patients who are sometimes asymptomatic carriers of infection.

Chlamydia trachomatis is responsible for nearly 30% of nongonococcal urethritis, or NGU. Chlamydia trachomatis urethritis may be discreet, the disease then progresses to a certain form of chronicity. The diagnosis will, like for the other clinical forms of the disease, be called into play later.

Chlamydia trachomatis is a cause of epididymitis in humans during a period of sexual activity. The bacterium may be found in the urethra, urine, sperm or even a sample collected by aspiration from the epididymis. It is in particular found in humans under 35 years of age. A discharge from the urethra which is associated with the disease suggests the diagnosis of a Chlamydia condition or sometimes a gonococcal condition.

Untreated Reiter's syndrome, if accompanied by urethritis, evokes a Chlamydia trachomatis condition.

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gonorrhoea (or have had contact), 10% to 20% of women having a venereal origin, 5% of women consulting having no particular origin.

The cervix is often normal during a Chlamydia trachomatis infection. However, a hypertrophic cervical erythema will cause such an infection to be suspected. Chlamydia trachomatis is responsible for an endocervicitis whereas viral impairments result in exocervicitis. A nongonococcal endocervicitis requires treating the patient and partners with tetracyclines.

Chlamydia trachomatis is responsible for a large number of acute salpingites. The picture is often complicated by an acute peritonitis or even a perihepatitis.

In case of pregnancy, the risk is first that of infection of the neonate at birth.

However, the risk of postpartum complications exists (endometritis or salpingitis).

The reference method for the diagnosis of *Chlamydia trachomatis* is the isolation of the bacterium on cell culture. For all infections, the sample collection should make it possible to obtain a suitable sample with the aid of a swab. This sample should be transported to a laboratory under excellent conditions; in particular, the cold chain must absolutely be maintained. The placing in cell culture on mouse fibroblasts will be carried out by people having specific skills. The distinction of *Chlamydia trachomatis* with labelled antibodies and the observation of cell cultures under a microscope will take place two days after placing in culture. Provided these imperatives are observed, cell culture is a reliable technique. However, the constraints linked to this technique are many: not only must the laboratory be equipped for the cell culture, but, furthermore, highly competent staff must take care of this type of diagnosis.

Techniques for identifying genetic material can obviously be used for the detection of Chlamydia trachomatis. Among these techniques, enzymatic gene amplification or PCR is favoured by those skilled in the art. The technique indeed makes it possible to identify Chlamydia trachomatis with a very high sensitivity and complete specificity. Initially used in specialist laboratories, PCR is now performed in numerous medical laboratories. This diagnostic approach is important because it allows detection of the bacteria even in samples which have been transported under poor conditions.

The treatment of *Chlamydia* urethritis with antibiotics such as tetracycline or quinolones is very effective. The duration of treatment varies between 7 and 14 days. The treatment of pregnant women poses the problem of contraindications to tetracycline.

Neonatal infections caused by *Chlamydia trachomatis* are explained by the frequency of these bacteria in the cervix. In some studies, 5% to 13% of impairments are observed in the cervix in asymptomatic pregnant women. The neonates risk, in this case, developing an inclusion conjunctivitis. Not only can *Chlamydia trachomatis* be isolated from the children's eyes, but also persistently from the rhinopharynx and also from the rectum. Pneumopathies and otitis media are also found, a result of contamination at childbirth.



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Differential diagnosis of inclusion conjunctivitis in neonates is required with gonococcal ophthalmia; while the duration of incubation is from one to three days in the case of a gonococcal ophthalmia, neonatal inclusion conjunctivitis has an acute beginning with discharge and formation of membranes or even of conjunctival scars.

Treatment consists of oral erythromycin at the dose of 40 to 50 mg per kg of weight, for two to three weeks. In a nonendemic trachoma region, this disease never progresses to chronicity.

Finally, mention should be made of infantile pneumopathy. The syndrome is well defined; it is found in children affected by *Chlamydia trachomatis*. Less than ten children are affected by *Chlamydia trachomatis* pneumopathies per thousand births. The syndrome is, in this case, always found at an early age (less than four months).

Venereal lymphogranulomatosis is an infection which is transmitted through sexual contact and is due to *Chlamydia trachomatis* strains L1, L2 and L3. In humans, a passing primary genital lesion is followed by an often suppurative and multiple regional lymphadenopathy. This disease is a general disease which is accompanied by fever and a rise in the number of white blood cells. If it progresses to chronicity, the disease then becomes complicated with genital elephantiasis, stricture or even fistula of the genital apparatus, of the penis, of the urethra and of the rectum.

Imphogranulomatosis. These Chlamydia strains are more virulent than the strains responsible for trachoma and STD. It is very important to note that venereal lymphogranulomatosis is a systemic disease which affects primarily the lymphatic tissue. Generally transmitted by the sexual route, Chlamydia trachomatis L may also cause contamination through direct contact or even during poor laboratory handling. In spite of these variable modes of transmission, the age for the highest incidence of these diseases corresponds to that for greater sexual activity. Venereal lymphogranulomatosis is still endemic in South America, in Africa and sometimes in Asia. For a long time, the prevalence of venereal lymphogranulomatosis was difficult to establish because of the difficulty of performing diagnosis with certitude. It should also be noted that men are affected more often than women. In low endemic regions, it is difficult to recognize the reservoir of microbes. This situation is explained by the fact that the isolation of the strains causing venereal lymphogranulomatoses from asymptomatic subjects is rarely successful.

Clinical impairment by venereal lymphogranulomatosis manifests itself by the appearance of a small ulcer 3 to 21 days after the exposure of small nonpainful vesicles. In both men and women, the lesion is most often silent. Since this impairment disappears within a few days and causes no functional discomfort and leaves no visible scar, the disease is often recognized late. The venereal lymphogranulomatosis strains may be found in the urethra or the endocervix in patients with inguinal adenopathies; these regions are then considered as the initial site of infection. The

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characteristic feature of the venereal lymphogranulomatosis strains is that from the initial site of infection, Chlamydia exhibits a diffusion drained by the lymphatic ducts. The disease is then complicated by a ganglionic impairment of the region draining the site of inoculation. By way of example, anorectal infection causes deep adenopathies. These adneopathies are marked by the appearance of a periadenitis which forms a fluctuating and suppurative ganglionic mass. Fistulae will appear during the decline of the disease. As general signs are present at this stage of the disease, it is often confused with a malignant lymphoma. The other general complications are rarely observed. Clinical examinations have been able to lead biologists to isolate Chlamydia from the cerebrospinal fluid or from the blood. It should also be noted that in a number of cases (5%), venereal lymphogranulomatosis is complicated by a chronic oedema: this is genital elephantiasis.

The diagnosis of venereal lymphogranulomatosis requires the isolation of the Chlamydia strains involved in the disease. However, isolation on cell cultures is rarely used, but immunological reactions may be used.

The treatment of venereal lymphogranulomatosis in its initial phase is identical to the treatment of other *Chlamydia* infections. In the chronic phases, antibiotics have little effect on the progress of the disease, but they are however useful in case of superinfection. Although the recommended therapeutic arsenal is identical, it is advisable to prolong the treatment for a period of at least four weeks. In addition to this treatment, reconstructive surgery may be useful in cases of urethral, penile or rectal strictures, as well as for the treatment of fistulae.

In conclusion, a short and effective treatment, without recurrences, and a well-tolerated treatment of *Chlamydia trachomatis* infections therefore remains desirable.

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An even greater need up until now relates to a diagnosis which is specific to each of the strains, which is sensitive, which can be carried out conveniently and rapidly, and which allows early detection of the infection.

No vaccine is currently available against *Chlamydia trachomatis*. The role of the immune defense in the physiology and pathology of the disease should probably be understood in order to develop satisfactory vaccines.

More detailed information relating to the biology of these strains, their interactions with their hosts, the associated phenomena of infectivity and those of escaping the immune defenses of the host in particular, and finally their involvement in the development of the these associated pathologies, will allow a better understanding of these mechanisms. In the light of the preceding text which shows in particular the limitations of the means of controlling *Chlamydia trachomatis* infection, it is therefore at present essential, on the one hand, to develop molecular tools, in particular from a better genetic knowledge of *Chlamydia trachomatis*, but also to develop new preventive and therapeutic treatments, new diagnostic methods and new vaccin strategies which are specific,

effective and tolerated. This is precisely the object of the present invention.

The subject of the present invention is the nucleotide sequence having the sequence SEQ ID No. 1 of the Chlamydia trachomatis LGV2 genome. However, the invention is not limited to SEQ ID No. 1, but encompasses genomes and nucleotides encoding polypeptides of strain variants, 5 polymorphisms, allelic variants, and mutants.

Thus, the subject of the present invention encompasses nucleotide sequences characterized in that they are chosen from:

- the nucleotide sequence of SEQ ID No. 1, a nucleotide sequence exhibiting at least 99.9% identity with the sequence SEQ ID No. 1, the nucleotide sequence of the genomic DNA 10 contained within ECACC Deposit No. 98112618, the nucleotide sequence of a clone insert within ECACC Deposit No. 98112617 (these being provisional deposit numbers);
  - a nucleotide sequence homologous to the sequence SEQ ID No. 1; b)
  - a polynucleotide sequence that hybridizes to the nucleotide sequence of a) under c) conditions of high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 µg/ml denatured salmon sperm DNA and 5-20 X 10<sup>6</sup> cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, 25 or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.
- (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and th hybridized probes are 35 detectable by autoradiography. Other conditions of intermediate stringency which may be used are

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well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.

- d) a nucleotide sequence complementary to the sequence SEQ ID No. 1 or complementary to a nucleotide sequence as defined in a), b) or c), and a nucleotide sequence of their corresponding RNA;
- e) a nucleotide sequence of a representative fragment of the sequence SEQ ID No. 1, or of a representative fragment of the nucleotide sequence as defined in a), b), c) or d);
  - f) a nucleotide sequence comprising a sequence as defined in a), b), c), d) or e);
- g) a nucleotide sequence capable of being obtained from a nucleotide sequence as defined in a), b), c), d), e) or f); and
- h) a modified nucleotide sequence of a nucleotide sequence as defined in a), b), c), d), e), f) or g).
- Sequence of the genome, or genomic sequence of *Chlamydia trachomatis* is understood to mean the sequence of the chromosome of *Chlamydia trachomatis*, in contrast with the plasmid sequence of *Chlamydia trachomatis*.

Nucleotide sequence, polynucleotide or nucleic acid are understood to mean, according to the present invention, either a double-stranded DNA, a single-stranded DNA or products of transcription of the said DNAs.

It should be understood that the present invention does not relate to the genomic nucleotide sequences of *Chlamydia trachomatis* taken in their natural environment, that is to say in the natural state. They are sequences which may have been isolated, purified or partially purified, by separation methods such as, for example, ion-exchange chromatography, molecular size exclusion chromatography or affinity chromatography, or alternatively fractionation techniques based on solubility in various solvents, or by genetic engineering methods such as amplification, cloning or subcloning, it being possible for the sequences of the invention to be carried by vectors.

The nucleotide sequence SEQ ID No. 1 was obtained by sequencing the *Chlamydia trachomatis LGV2* genome by the method of directed sequencing after fluorescent automated sequencing of the inserts of clones and assembling of these sequences of nucleotide fragments (inserts) by means of softwares (cf. Examples). In spite of the high precision of the sequence SEQ ID No. 1, it is possible that it does not perfectly, 100% represent the nucleotide sequence of the *Chlamydia trachomatis LGV2* genome and that a few rare sequencing errors or uncertainties still remain in the sequence SEQ ID No. 1. In the present invention, the presence of an uncertainty for an amino acid is designated by «Xaa» and that for a nucleotide is designated by «N» in the sequence

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listing below. These few rare errors or uncertainties could be easily detected and corrected by persons skilled in the art using the entire chromosome and/or its representative fragments according to the invention and standard amplification, cloning and sequencing methods, it being possible for the sequences obtained to be easily compared, in particular by means of a computer software and using computer-readable media for recording the sequences according to the invention as described, for example, below. After correcting these possible rare errors or uncertainties, the corrected nucleotide sequence obtained would still exhibit at least 99.9% identity with the sequence SEQ ID No. 1. Such rare sequencing uncertainties are not present within the DNA contained within ECACC Deposit No. 98112617 or 98112618 (provisional numbers) and whatever rare sequence uncertainties that exist within SEQ ID No. 1 can routinely be corrected utilizing the DNA of the ECACC Deposits.

Homologous nucleotide sequence for the purposes of the present invention is understood to mean a nucleotide sequence having a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, this percentage being purely statistical and it being possible for the differences between the two nucleotide sequences to be 15 distributed randomly and over their entire length. The said homologous sequences exhibiting a percentage identity with the bases of the nucleotide sequence SEQ ID No. 1 of at least 80%, preferably 90% and 95%, may comprise, for example, the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the Chlamydia family, including the species Chlamydia pneumoniae, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the genomic sequence or to the sequences of its representative fragments of a bacterium belonging to the variants of the species Chlamydia trachomatis. In the present invention, the terms family and genus are mutually interchangeable, the terms variant, serotype, strain and subspecies are also mutually interchangeable. These homologous sequences may thus correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in Chlamydia.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

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In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well known in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

- (1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence database;
  - (2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database;
- (3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;
- (4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and
- (5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g., Karlin and Altschul, 1990, *Proc. Natl. Acad. Sci. USA 87*:2267-2268).

Nucleotide sequence complementary to a sequence of the invention is understood to mean any DNA whose nucleotides are complementary to those of the sequence of the invention, and whose orientation is reversed (antiparallel sequence).

The present invention further comprises fragments of the sequences of a) through h) above. Representative fragments of the sequences according to the invention will be understood to mean any nucleotide fragment having at least 8 successive nucleotides, preferably at least 12

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successive nucleotides, and still more preferably at least 15 or at least 20 successive nucleotides of the sequence from which it is derived. It is understood that such fragments refer only to portions of SEQ ID No. 1 that are not currently listed in a publicly available database.

Among these representative fragments, those capable of hybridizing under stringent conditions with a nucleotide sequence according to the invention are preferred. Hybridization under stringent conditions means that the temperature and ionic strength conditions are chosen such that they allow hybridization to be maintained between two complementary DNA fragments.

By way of illustration, high stringency conditions for the hybridization step for the purposes of defining the nucleotide fragments described above, are advantageously the following.

The hybridization is carried out at a preferred temperature of 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15 M NaCl and 0.05 M Na citrate. The washing steps may be, for example, the following:

2 x SSC, 0.1% SDS at room temperature followed by three washes with 1 x SSC, 0.1% SDS; 0.5 x SSC, 0.1% SDS; 0.1 x SSC, 0.1% SDS at 68°C for 15 minutes.

Intermediate stringency conditions, using, for example, a temperature of 60°C in the presence of a 5 x SSC buffer, or of low stringency, for example a temperature of 50°C in the presence of a 5 x SSC buffer, respectively require a lower overall complementarity for the hybridization between the two sequences.

The stringent hybridization conditions described above for a polynucleotide of about 300 bases in size will be adapted by persons skilled in the art for larger- or smaller-sized oligonucleotides, according to the teaching of Sambrook et al., 1989.

Among the representative fragments according to the invention, those which can be used as primer or probe in methods which make it possible to obtain homologous sequences or their representative fragments according to the invention, or to reconstitute a genomic fragment found to be incomplete in the sequence SEQ ID No. 1 or carrying an error or an uncertainty, are also preferred, these methods, such as the polymerase chain reaction (PCR), cloning and sequencing of nucleic acid being well known to persons skilled in the art. These homologous nucleotide sequences corresponding to mutations or to inter- or intra-species variations, as well as the complete genomic sequence or one of its representative fragments capable of being reconstituted, of course form part of the invention.

Among the said representative fragments, those which can be used as primer or probe in methods allowing diagnosis of the presence of *Chlamydia trachomatis* or one of its associated microorganisms as defined below are also preferred.

The representative fragments capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia trachomatis* or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia trachomatis* or one of its associated

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microorganisms in the host cell and/or organism, are also preferred. Replication cycle is intended to designate invasion, multiplication, intracellular localization, in particular retention in the vacuole and inhibition of the process of fusion to the lysosome, and propagation of *Chlamydia trachomatis* or one

of its associated microorganisms from host cells to host cells.

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Among the said representative fragments, those corresponding to nucleotide sequences corresponding to open reading frames, called ORF sequences (ORF for open reading frame), and encoding polypeptides, such as for example, but without being limited thereto, the ORF sequences which will be later described, are finally preferred.

The representative fragments according to the invention may be obtained, for example, by specific amplification, such as PCR, or after digestion, with appropriate restriction enzymes, of nucleotide sequences according to the invention; these methods are in particular described in the manual by Sambrook et al., 1989. The said representative fragments may also be obtained by chemical synthesis when they are not too large in size and according to methods well known to persons skilled in the art. For example, such fragments can be obtained by isolating fragments of the genomic DNA of ECACC Deposit No. 98112618 or a clone insert present at this ECACC Deposit No. 98112617 (provisional numbers).

The representative fragments according to the invention may be used, for example, as primer, to reconstitute some of the said representative fragments, in particular those in which a portion of the sequence is likely to be missing or imperfect, by methods well known to persons skilled in the art such as amplification, cloning or sequencing techniques.

Modified nucleotide sequence will be understood to mean any nucleotide sequence obtained by mutagenesis according to techniques well known to persons skilled in the art, and exhibiting modifications in relation to the normal sequences, for example mutations in the regulatory and/or promoter sequences for the expression of a polypeptide, in particular leading to a modification of the level of expression of the said polypeptide or to a modulation of the replicative cycle.

Modified nucleotide sequence will also be understood to mean any nucleotide sequence encoding a modified polypeptide as defined below.

The subject of the present invention also includes *Chlamydia trachomatis* nucleotide sequences characterized in that they are chosen from a nucleotide sequence of an open reading frame (ORF), that is, the ORF2 to ORF1197 sequences.

The ORF2 to ORF1197 nucleotide sequences are defined in Tables 1 and 2, infra, represented below by their position on the sequence SEQ ID No. 1. For example, the ORF10 sequence is defined by the nucleotide sequence between the nucleotides at position 9828 and 10430 on the sequence SEQ ID No. 1, ends included. ORF2 to ORF1197 have been identified via homology analyses as well as via analyses of potential ORF start sites, as discussed in the examples

below. It is to be understood that each identified ORF of the invention comprises a nucleotide sequence that spans the contiguous nucleotide sequence from the codon immediately 3' to the stop codon of the preceding ORF and through the 5' codon to the next stop codon of SEQ ID No.:1 inframe to the ORF nucleotide sequence. Table 2, *infra*, lists the beginning, end and potential start site of each of ORFs 2-1197. In one embodiment, the ORF comprises the contiguous nucleotide sequence spanning from the potential ORF start site downstream (that is, 3') to the ORF stop codon (or the ORF codon immediately adjacent to and upstream of the ORF stop codon). ORF2 to ORF1197 encode the polypeptides of SEQ ID No. 2 to SEQ ID No. 1197.

Upon introduction of minor frameshifts, certain individual ORFs can comprise larger combined ORFs. A list of such putative combined ORFs is shown in Table 3, below. For example, a combined ORF can comprise ORF 1076 and ORF 1073, including intervening in-frame, nucleotide sequences. The order of ORFs (5' to 3'), within each combined ORF is as listed. It is to be understood that when ORF2 to ORF1197 are referred to herein, such reference is also meant to include combined ORFs. Polypeptide sequences encoded by such combined ORFs are also part of the present invention.

#### Table 3

ORF 1076, ORF 1073;

ORF 3, ORF 2;

ORF 23, ORF 22, ORF 21;

20 ORF 1141, ORF 477, ORF 478, ORF 479;

ORF 487, ORF 486, ORF 485, ORF 484, ORF 483, ORF 482, ORF 481;

ORF 488, ORF 489;

ORF 573, ORF 572, ORF 571:

ORF 817, ORF 818;

25 ORF 819, ORF 820;

ORF 1037, ORF 1038;

ORF 1071, ORF 1070;

ORF 17, ORF 1077;

ORF 1185, ORF 933, ORF 934;

30 ORF 1060, ORF 1059;

ORF 155, ORF 156;

ORF 679, ORF 680;

ORF 879, ORF 878;

ORF 1028; ORF 1029,

35 and representative fragments.

Table 1 also depicts the results of homology searches that compared the sequences of the polypeptides encoded by each of the ORFs to sequences present in public published databases. It is understood that in one embodiment, those polypeptides listed in Table 1 as exhibiting greater than about 95% identity to a polypeptide present in a publicly disclosed database are not considered part of the present invention; likewise in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention. In another embodiment, it is understood that those polypeptides listed in Table 1 as exhibiting greater than about 99% identity to a polypeptide present in a publicly disclosed database are not considered part of the invention; likewise, in this embodiment, those nucleotide sequences encoding such polypeptides are not considered part of the invention.

The invention also relates to the nucleotide sequences characterized in that they comprise a nucleotide sequence chosen from:

- a) an ORF2 to ORF1197, a «combined» ORF nucleotide sequence, the nucleotide sequence of the genomic DNA contained within ECACC Deposit No. 98112618 or the nucleotide
   sequence of a clone insert in ECACC Deposit No. 98112617 according to the invention;
  - b) a homologous nucleotide sequence exhibiting at least 80% identity across an entire ORF2 to ORF1197 nucleotide sequence according to the invention or as defined in a);
  - c) a polynucleotide sequence that hybridizes to ORF2 to ORF1197 under conditions of high or intermediate stringency as described below:
- 20 (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100  $\mu g/ml$  denatured salmon sperm DNA and 5-20 X 10<sup>6</sup> cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x 30 SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety.

Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a *Chlamydia trachomatis* polypeptide.

- (ii) By way of example and not limitation, procedures using conditions of intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.
- d) a complementary or RNA nucleotide sequence corresponding to an ORF2 to ORF1197 sequence according to the invention or as defined in a), b) or c);
  - e) a nucleotide sequence of a representative fragment of an ORF2 to ORF1197 sequence according to the invention or of a sequence as defined in a), b), c) or d);
  - f) a nucleotide sequence capable of being obtained from an ORF2 to ORF1197 sequence according to the invention or as defined in a), b), c), d) or e); and
- g) a modified nucleotide sequence of an ORF2 to ORF1197 sequence according to the invention or as defined in a), b), c), d), e) or f).

As regards the homology with the ORF2 to ORF1197 nucleotide sequences, the homologous sequences exhibiting a percentage identity with the bases of one of the ORF2 to ORF1197 nucleotide sequences of at least 80%, preferably 90% and 95%, are preferred. Such homologous sequences are identified routinely via, for example, the algorithms described above and in the examples below. The said homologous sequences correspond to the homologous sequences as defined above and may comprise, for example, the sequences corresponding to the ORF sequences of a bacterium belonging to the Chlamydia family, including the species Chlamydia pneumoniae, Chlamydia psittaci and Chlamydia pecorum mentioned above, as well as the sequences corresponding to the ORF sequences of a bacterium belonging to the variants of the species Chlamydia trachomatis. These homologous sequences may likewise correspond to variations linked to mutations within the same species or between species and may correspond in particular to truncations, substitutions, deletions and/or additions of at least one nucleotide. The said homologous sequences may also correspond to variations linked to the degeneracy of the genetic code or to a bias in the genetic code which is specific to the family, to the species or to the variant and which are likely to be present in

Chlamydia.

The invention comprises the polypeptides encoded by a nucleotide sequence according to the invention, preferably by a representative fragment of the sequence SEQ ID No. 1 and corresponding to an ORF sequence, in particular the *Chlamydia trachomatis* polypeptides, characterized in that they are chosen from the sequences SEQ ID No. 2 to SEQ ID No. 1197, and representative fragments thereof. However, the invention is not limited to polypeptides encoded by ORFs in SEQ ID No. 1 and its corresponding ORF sequences, but encompasses polypeptides of strain variants, polymorphisms, allelic variants, and mutants.

Thus, the invention also comprises the polypeptides characterized in that they comprise a polypeptide chosen from:

- a) a polypeptide encoded by a polynucleotide sequence in SEQ ID No. 1 (e.g., any polypeptide encoded by a polynucleotide sequence corresponding to ORF2 to ORF1197) and/or representative fragments thereof according to the invention;
- b) a polypeptide homologous to a polypeptide according to the invention, or as defined 15 in a);
  - c) a polypeptide encoded by a polynucleotide sequence that hybridizes to SEQ ID No. 1 or ORF2 to ORF1197 under high or intermediate stringency as described below:
- (i) By way of example and not limitation, procedures using conditions of high stringency are as follows: Prehybridization of filters containing DNA is carried out for 8 h to 20 overnight at 65°C in buffer composed of 6X SSC, 50 mM Tris-HCl (pH 7.5), 1 mM EDTA, 0.02% PVP, 0.02% Ficoll, 0.02% BSA, and 500 µg/ml denatured salmon sperm DNA. Filters are hybridized for 48 h at 65°C, the preferred hybridization temperature, in prehybridization mixture containing 100 μg/ml denatured salmon sperm DNA and 5-20 X 106 cpm of <sup>32</sup>P-labeled probe. Alternatively, the hybridization step can be performed at 65°C in the presence of SSC buffer, 1 x SSC corresponding to 25 0.15M NaCl and 0.05 M Na citrate. Subsequently, filter washes can be done at 37°C for 1 h in a solution containing 2X SSC, 0.01% PVP, 0.01% Ficoll, and 0.01% BSA, followed by a wash in 0.1X SSC at 50°C for 45 min. Alternatively, filter washes can be performed in a solution containing 2 x SSC and 0.1% SDS, or 0.5 x SSC and 0.1% SDS, or 0.1 x SSC and 0.1% SDS at 68°C for 15 minute intervals. Following the wash steps, the hybridized probes are detectable by autoradiography. Other 30 conditions of high stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a polypeptide encoded by one of ORF2 to ORF1197 . In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.

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- By way of example and not limitation, procedures using conditions of (ii) intermediate stringency are as follows: Filters containing DNA are prehybridized, and then hybridized at a temperature of 60°C in the presence of a 5 x SSC buffer and labeled probe. Subsequently, filters washes are performed in a solution containing 2x SSC at 50°C and the hybridized probes are detectable by autoradiography. Other conditions of intermediate stringency which may be used are well known in the art and as cited in Sambrook et al., 1989, Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Press, N.Y., pp. 9.47-9.57; and Ausubel et al., 1989, Current Protocols in Molecular Biology, Green Publishing Associates and Wiley Interscience, N.Y. are incorporated herein in their entirety. Preferably, such sequences encode a homolog of a 10 polypeptide encoded by one of ORF2 to ORF1197. In one embodiment, such sequences encode a Chlamydia trachomatis polypeptide.
  - a fragment of at least 5 amino acids of a polypeptide according to the invention, or as d) defined in a), b) or c);
- a biologically active fragment of a polypeptide according to the invention, or as 15 defined in a), b), c) or d); and
  - f) a modified polypeptide of a polypeptide according to the invention, as defined in a), b), c), d) or e).

In the present description, the terms polypeptide, peptide and protein are interchangeable.

20 It should be understood that the invention does not relate to the polypeptides in natural form, that is to say that they are not taken in their natural environment but that they may have been isolated or obtained by purification from natural sources, or alternatively obtained by genetic recombination, or else by chemical synthesis and that they may, in this case, comprise nonnatural amino acids, as will be described below.

Homologous polypeptide will be understood to designate the polypeptides exhibiting, in relation to the natural polypeptide, certain modifications such as in particular a deletion, addition or substitution of at least one amino acid, a truncation, an extension, a chimeric fusion, and/or a mutation, or polypeptides exhibiting post-translational modifications. Among the homologous polypeptides, those whose amino acid sequence exhibits at least 80%, preferably 90%, homology or 30 identity with the amino acid sequences of the polypeptides according to the invention are preferred. In the case of a substitution, one or more consecutive or nonconsecutive amino acids are replaced by «equivalent» amino acids. The expression «equivalent» amino acid is intended here to designate any amino acid capable of being substituted for one of the amino acids in the basic structure without, however, essentially modifying the biological activities of the corresponding peptides and as will be defined later.

Protein and/or nucleic acid sequence homologies may be evaluated using any of the variety of sequence comparison algorithms and programs known in the art. Such algorithms and programs include, but are by no means limited to, TBLASTN, BLASTP, FASTA, TFASTA, and CLUSTALW (Pearson and Lipman, 1988, Proc. Natl. Acad. Sci. USA 85(8):2444-2448; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Thompson et al., 1994, Nucleic Acids Res. 22(2):4673-4680; Higgins et al., 1996, Methods Enzymol. 266:383-402; Altschul et al., 1990, J. Mol. Biol. 215(3):403-410; Altschul et al., 1993, Nature Genetics 3:266-272).

In a particularly preferred embodiment, protein and nucleic acid sequence homologies are evaluated using the Basic Local Alignment Search Tool ("BLAST") which is well know in the art (see, e.g., Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268; Altschul et al., 1990, J. Mol. Biol. 215:403-410; Altschul et al., 1993, Nature Genetics 3:266-272; Altschul et al., 1997, Nuc. Acids Res. 25:3389-3402). In particular, five specific BLAST programs are used to perform the following task:

(1)BLASTP and BLAST3 compare an amino acid query sequence against a protein sequence 15 database;

(2)BLASTN compares a nucleotide query sequence against a nucleotide sequence database;

(3)BLASTX compares the six-frame conceptual translation products of a query nucleotide sequence (both strands) against a protein sequence database;

(4)TBLASTN compares a query protein sequence against a nucleotide sequence database translated in all six reading frames (both strands); and

(5)TBLASTX compares the six-frame translations of a nucleotide query sequence against the six-frame translations of a nucleotide sequence database.

The BLAST programs identify homologous sequences by identifying similar segments, which are referred to herein as "high-scoring segment pairs," between a query amino or nucleic acid sequence and a test sequence which is preferably obtained from a protein or nucleic acid sequence database. High-scoring segment pairs are preferably identified (i.e., aligned) by means of a scoring matrix, many of which are known in the art. Preferably, the scoring matrix used is the BLOSUM62 matrix (Gonnet et al., 1992, Science 256:1443-1445; Henikoff and Henikoff, 1993, Proteins 17:49-61). Less preferably, the PAM or PAM250 matrices may also be used (see, e.g., Schwartz and Dayhoff, eds., 1978, Matrices for Detecting Distance Relationships: Atlas of Protein Sequence and Structure, Washington: National Biomedical Research Foundation)

The BLAST programs evaluate the statistical significance of all high-scoring segment pairs identified, and preferably selects those segments which satisfy a user-specified threshold of significance, such as a user-specified percent homology. Preferably, the statistical significance of a high-scoring segment pair is evaluated using the statistical significance formula of Karlin (see, e.g.,

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Karlin and Altschul, 1990, Proc. Natl. Acad. Sci. USA 87:2267-2268).

Equivalent amino acids may be determined either based on their structural homology with the amino acids for which they are substituted, or on results of comparative tests of biological activity between the various polypeptides which may be carried out.

By way of example, there may be mentioned the possibilities of substitutions which may be carried out without resulting in a substantial modification of the biological activity of the corresponding modified polypeptides; the replacements, for example, of leucine with valine or isoleucine, of aspartic acid with glutamic acid, of glutamine with asparagine, of arginine with lysine, and the like, the reverse substitutions naturally being feasible under the same conditions.

The homologous polypeptides also correspond to the polypeptides encoded by the homologous nucleotide sequences as defined above and thus comprise in the present definition the mutated polypeptides or polypeptides corresponding to inter- or intra-species variations which may exist in *Chlamydia*, and which correspond in particular to truncations, substitutions, deletions and/or additions of at least one amino acid residue.

Biologically active fragment of a polypeptide according to the invention will be understood to designate in particular a polypeptide fragment, as defined below, exhibiting at least one of the characteristics of the polypeptides according to the invention, in particular in that it is:

- capable of eliciting an immune response directed against Chlamydia trachomatis; and/or
- capable of being recognized by an antibody specific for a polypeptide according to the invention; and/or
- capable of binding to a polypeptide or to a nucleotide sequence of *Chlamydia trachomatis*; and/or
- capable of modulating, regulating, inducing or inhibiting the expression of a gene of *Chlamydia trachomatis* or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia trachomatis* or one of its associated microorganisms in the host cell and/or organism; and/or
- capable of generally exerting an even partial physiological activity, such as for example a structural activity (cellular envelope, ribosome), an enzymatic (metabolic) activity, a transport activity, an activity in the secretion or in the virulence.

A representative polypeptide fragment according to the invention is understood to designate a polypeptide comprising a minimum of 5 amino acids, preferably 10 amino acids or preferably 15 amino acids. It is to be understood that such fragments refer only to portions of polypeptides encoded by ORF2 or ORF1197 that are not currently listed in a publicly available database.

The polypeptide fragments according to the invention may correspond to isolated or

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purified fragments which are naturally present in Chlamydia trachomatis or which are secreted by Chlamydia trachomatis, or may correspond to fragments capable of being obtained by cleaving the said polypeptide with a proteolytic enzyme, such as trypsin or chymotrypsin or collagenase, or with a chemical reagent, such as cyanogen bromide (CNBr) or alternatively by placing the said polypeptide in a highly acidic environment, for example at pH 2.5. Such polypeptide fragments may be equally well prepared by chemical synthesis, using hosts transformed with an expression vector according to the invention containing a nucleic acid allowing the expression of the said fragments, placed under the control of appropriate elements for regulation and/or expression.

«Modified polypeptide» of a polypeptide according to the invention is understood to designate a polypeptide obtained by genetic recombination or by chemical synthesis as will be described below, exhibiting at least one modification in relation to the normal sequence. These modifications may in particular affect amino acids responsible for a specificity or for the efficiency of the activity, or responsible for the structural conformation, for the charge or for the hydrophobicity, and for the capacity for multimerization and for membrane insertion of the polypeptide according to the invention. It is thus possible to create polypeptides with an equivalent, an increased or a reduced activity, and with an equivalent, a narrower or a broader specificity. Among the modified polypeptides, there may be mentioned the polypeptides in which up to 5 amino acids may be modified, truncated at the N- or C-terminal end, or alternatively deleted, or else added.

As is indicated, the modifications of the polypeptide may have in particular the 20 objective:

- of making it capable of modulating, regulating, inhibiting or inducing the expression of a gene of *Chlamydia*, in particular of *Chlamydia trachomatis* and its variants, or one of its associated microorganisms, and/or capable of modulating the replication cycle of *Chlamydia*, in particular of *Chlamydia trachomatis* and its variants, or one of its associated microorganisms, in the host cell and/or organism,
- of allowing its use in methods of biosynthesis or of biodegradation, or its incorporation into vaccine compositions,
  - of modifying its bioavailability as a compound for therapeutic use.

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The said modified polypeptides may also be used on any cell or microorganism for which the said modified polypeptides will be capable of modulating, regulating, inhibiting or inducing gene expression, or of modulating the growth or the replication cycle of the said cell or of the said microorganism. The methods allowing demonstration of the said modulations on eukaryotic or prokaryotic cells are well known to persons skilled in the art. The said cells or microorganisms will be chosen, in particular, from tumour cells or infectious microorganisms and the said modified polypeptides may be used for the prevention or treatment of pathologies linked to the presence of the

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said cells or of the said microorganisms. It is also clearly understood that the nucleotide sequences encoding the said modified polypeptides may be used for the said modulations, for example by the intermediacy of vectors according to the invention and which are described below, so as to prevent or to treat the said pathologies.

The above modified polypeptides may be obtained using combinatory chemistry, in which it is possible to systematically vary portions of the polypeptide before testing them on models, cell cultures or microorganisms for example, so as to select the compounds which are the most active or which exhibit the desired properties.

Chemical synthesis also has the advantage of being able to use:

- nonnatural amino acids, or
  - nonpeptide bonds.

Accordingly, in order to extend the life of the polypeptides according to the invention, it may be advantageous to use nonnatural amino acids, for example in the D form, or alternatively amino acid analogues, in particular sulphur-containing forms for example.

Finally, the structure of the polypeptides according to the invention, its homologous or modified forms, as well as the corresponding fragments may be integrated into chemical structures of the polypeptide type and the like. Accordingly, it may be advantageous to provide at the N- and C-terminal ends compounds which are not recognized by proteases.

Also forming part of the invention are the nucleotide sequences encoding a 20 polypeptide according to the invention. Described below are ORF nucleotide sequences encoding polypeptides exhibiting particularly preferable characteristics. For each group of preferred ORFs described below, it is to be understood that in addition to the individual ORFs listed, in instances wherein such ORFs are present as part of «combined» ORFs, the «combined» ORFs are also to be included within the preferred group.

More particularly, the subject of the invention is nucleotide sequences, characterized in that they encode a polypeptide of the cellular envelope, preferably of the outer cellular envelope of Chlamydia trachomatis or one of its representative fragments, such as for example the predominant proteins of the outer membrane, the adhesion proteins or the proteins entering into the composition of the Chlamydia wall. Among these sequences, the sequences comprising a nucleotide sequence chosen 30 from the following sequences are most preferred:

ORF3; ORF19; ORF51; ORF189; ORF212; ORF213; ORF324; ORF477; ORF478; ORF479; ORF481; ORF482; ORF483; ORF484; ORF486; ORF488; ORF489; ORF490; ORF572; ORF573; ORF742; ORF817; ORF818; ORF820; ORF1035; ORF1036; ORF1037; ORF1038; ORF1070; ORF1071; ORF1073 and one of their representative fragments.

The structure of the cytoplasmic membranes and of the wall of bacteria is dependent

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on the associated proteins. The structure of the cytoplasmic membrane makes it impermeable to water, to water-soluble substances and to small-sized molecules (ions, small inorganic molecules, peptides or proteins). To enter into or to interfere with a cell or a bacterium, a ligand must establish a special relationship with a protein anchored in the cytoplasmic membrane (the receptor). These proteins which are anchored on the membrane play an important role in metabolism since they control the exchanges in the bacterium. These exchanges apply to molecules of interest for the bacterium (small molecules such as sugars and small peptides) as well as undesirable molecules for the bacterium such as antibiotics or heavy metals.

The double lipid layer structure of the membrane requires the proteins which are inserted therein to have hydrophobic domains of about twenty amino acids forming an alpha helix. Predominantly hydrophobic and potentially transmembrane regions may be predicted from the primary sequence of the proteins, itself deduced from the nucleotide sequence. The presence of one or more putative transmembrane domains raises the possibility for a protein to be associated with the cytoplasmic membrane and to be able to play an important metabolic role therein or alternatively for the protein thus exposed to be able to exhibit potentially protective epitopes.

If the proteins inserted into the membrane exhibit several transmembrane domains capable of interacting with one another via electrostatic bonds, it then becomes possible for these proteins to form pores which go across the membrane which becomes permeable for a number of substances. It should be noted that proteins which do not have transmembrane domains may also be anchored by the intermediacy of fatty acids in the cytoplasmic membrane, it being possible for the breaking of the bond between the protein and its anchor in some cases to be responsible for the release of the peptide outside the bacterium.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF2; ORF3; ORF5; ORF8; ORF9; ORF10; ORF11; ORF12; ORF17; ORF21; ORF26; ORF27; ORF28; ORF29; ORF30; ORF31; ORF33; ORF35; ORF37; ORF39; ORF40; ORF41; ORF42; ORF43; ORF44; ORF45; ORF46; ORF47; ORF48; ORF49; ORF52; ORF53; ORF55; ORF56; ORF58; ORF65; ORF66; ORF68; ORF70; ORF74; ORF75; ORF76; ORF78; ORF79; ORF81; ORF82; ORF83; ORF86; ORF91; ORF92; ORF94; ORF97; ORF100; ORF102; ORF103; ORF105; ORF106; ORF107; ORF109; ORF110; ORF111; ORF112; ORF113; ORF114; ORF115; ORF116; ORF117; ORF120; ORF122; ORF123; ORF130; ORF134; ORF135; ORF137; ORF140; ORF141; ORF143; ORF144; ORF145; ORF147; ORF148; ORF149; ORF150; ORF151; ORF155; ORF156; ORF162; ORF163; ORF164; ORF165; ORF166; ORF167; ORF168; ORF169; ORF170; ORF171;

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ORF173; ORF176; ORF177; ORF181; ORF183; ORF184; ORF186; ORF187; ORF188;
    ORF190; ORF191; ORF192; ORF194; ORF195; ORF196; ORF197; ORF198; ORF199; ORF201;
    ORF202; ORF204; ORF206; ORF207; ORF209; ORF212; ORF213; ORF217; ORF219; ORF220;
    ORF221; ORF222; ORF223; ORF224; ORF225; ORF227; ORF228; ORF231; ORF232; ORF234;
 5 ORF236; ORF237; ORF243; ORF244; ORF245; ORF247; ORF248; ORF249; ORF252; ORF254;
    ORF257; ORF260; ORF261; ORF263; ORF265; ORF266; ORF267; ORF270; ORF271; ORF272;
    ORF274; ORF276; ORF277; ORF278; ORF279; ORF282; ORF283; ORF284; ORF285; ORF287;
    ORF289; ORF290; ORF291; ORF294; ORF298; ORF305; ORF306; ORF310; ORF311; ORF313;
    ORF315; ORF316; ORF319; ORF320; ORF322; ORF323; ORF325; ORF326; ORF327; ORF328;
10 ORF330; ORF331; ORF332; ORF333; ORF334; ORF335; ORF336; ORF338; ORF339; ORF340;
    ORF341; ORF344; ORF345; ORF348; ORF349; ORF350; ORF351; ORF352; ORF353; ORF356;
    ORF357; ORF361; ORF362; ORF366; ORF367; ORF370; ORF372; ORF373;
    ORF375; ORF377; ORF378; ORF380; ORF382; ORF383; ORF384; ORF385; ORF387;
    ORF389; ORF390; ORF391; ORF393; ORF396; ORF398; ORF399; ORF403; ORF404; ORF406;
15 ORF407; ORF413; ORF414; ORF417; ORF418; ORF420; ORF421; ORF424; ORF426; ORF427;
    ORF428; ORF430; ORF433; ORF434; ORF435; ORF436; ORF437; ORF440; ORF443; ORF446;
    ORF448; ORF450; ORF451; ORF454; ORF455; ORF457; ORF458; ORF459; ORF463; ORF464;
    ORF466; ORF467; ORF468; ORF469; ORF470; ORF473; ORF474; ORF475; ORF476; ORF477;
    ORF479; ORF480; ORF481; ORF483; ORF484; ORF485; ORF486; ORF487; ORF488; ORF491;
20 ORF493; ORF496; ORF497; ORF498; ORF500; ORF501; ORF503; ORF504; ORF508; ORF512;
    ORF513; ORF514; ORF519; ORF521; ORF523; ORF524; ORF526; ORF527; ORF529; ORF530;
    ORF531; ORF532; ORF534; ORF536; ORF537; ORF538; ORF540; ORF541; ORF542; ORF543;
    ORF544; ORF545; ORF546; ORF5547; ORF551; ORF5552; ORF5553; ORF5558; ORF5559;
    ORF560; ORF561; ORF562; ORF566; ORF567; ORF568; ORF569; ORF571; ORF572; ORF574;
25 ORF575; ORF580; ORF582; ORF585; ORF587; ORF589; ORF592; ORF593; ORF595;
    ORF596; ORF599; ORF601; ORF602; ORF603; ORF604; ORF608; ORF609; ORF610;
    ORF611; ORF615; ORF616; ORF617; ORF618; ORF621; ORF622; ORF623; ORF624; ORF625;
    ORF628; ORF632; ORF633; ORF634; ORF635; ORF637; ORF638; ORF640; ORF641; ORF643;
    ORF646; ORF649; ORF651; ORF652; ORF653; ORF655; ORF658; ORF664;
30 ORF665; ORF666; ORF668; ORF669; ORF670; ORF671; ORF672; ORF673; ORF674; ORF676;
    ORF677; ORF688; ORF682; ORF683; ORF684; ORF686; ORF688; ORF689; ORF690;
    ORF691; ORF693; ORF695; ORF696; ORF698; ORF701; ORF703; ORF704; ORF705;
    ORF706; ORF707; ORF709; ORF710; ORF711; ORF712; ORF713; ORF714; ORF715; ORF717;
   ORF718; ORF720; ORF721; ORF722; ORF724; ORF726; ORF728; ORF729; ORF730; ORF731;
35 ORF732; ORF734; ORF737; ORF738; ORF739; ORF740; ORF742; ORF743; ORF744;
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ORF745; ORF746; ORF748; ORF750; ORF751; ORF752; ORF753; ORF754; ORF755; ORF757; ORF758; ORF760; ORF764; ORF766; ORF768; ORF769; ORF771; ORF772; ORF773; ORF774; ORF775; ORF776; ORF777; ORF778; ORF780; ORF781; ORF782; ORF783; ORF786; ORF787; ORF788; ORF789; ORF790; ORF793; ORF798; ORF800; ORF802; ORF803; 5 ORF806; ORF808; ORF809; ORF810; ORF811; ORF813; ORF814; ORF817; ORF820; ORF822; ORF824; ORF825; ORF827; ORF828; ORF829; ORF830; ORF833; ORF834; ORF835; ORF837; ORF838; ORF839; ORF840; ORF841; ORF842; ORF843; ORF845; ORF848; ORF849; ORF850; ORF851; ORF852; ORF854; ORF855; ORF856; ORF857; ORF859; ORF860; ORF862; ORF863; ORF864; ORF866; ORF869; ORF872; ORF873; ORF874; ORF878; ORF879; ORF880; ORF881; 10 ORF883; ORF884; ORF885; ORF886; ORF887; ORF892; ORF893; ORF894; ORF895; ORF897; ORF899; ORF900; ORF901; ORF904; ORF906; ORF909; ORF910; ORF912; ORF914; ORF917; ORF920; ORF921; ORF922; ORF923; ORF924; ORF925; ORF926; ORF927; ORF930; ORF933; ORF934; ORF935; ORF936; ORF937; ORF940; ORF941; ORF942; ORF943; ORF944; ORF945; ORF947; ORF951; ORF952; ORF953; ORF954; ORF955; ORF956; ORF957; ORF958; 15 ORF960; ORF961; ORF962; ORF963; ORF964; ORF966; ORF967; ORF969; ORF970; ORF971; ORF973; ORF974; ORF979; ORF980; ORF981; ORF982; ORF984; ORF988; ORF989; ORF990; ORF991; ORF995; ORF999; ORF1001; ORF1003; ORF1004; ORF1005; ORF1006; ORF1007; ORF1009; ORF1010; ORF1011; ORF1012; ORF1013; ORF1014; ORF1016; ORF1017; ORF1018; ORF1020; ORF1021; ORF1025; ORF1026; ORF1027; ORF1029; ORF1030; ORF1031; 20 ORF1035; ORF1036; ORF1037; ORF1038; ORF1039; ORF1040; ORF1044; ORF1045; ORF1047; ORF1048; ORF1050; ORF1051; ORF1052; ORF1053; ORF1055; ORF1056; ORF1057; ORF1058; ORF1061; ORF1062; ORF1063; ORF1064; ORF1065; ORF1066; ORF1068; ORF1069; ORF1072; ORF1074; ORF1076 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 4 and 6 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF7; ORF14; ORF16; ORF32; ORF34; ORF36; ORF38; ORF50; ORF57; ORF59; ORF61; ORF62; ORF63; ORF64; ORF67; ORF69; ORF72; ORF77; ORF80; ORF84; ORF87; ORF93; ORF95; ORF99; ORF108; ORF119; ORF125; ORF126; ORF129; ORF131; ORF136; ORF139; ORF146; ORF152; ORF154; ORF160; ORF161; ORF172; ORF179; ORF182; ORF185; ORF200; ORF203; ORF205; ORF239; ORF242; ORF250; ORF253; ORF256; ORF259; ORF262; ORF268; ORF275; ORF281; ORF286; ORF288; ORF292; ORF295; ORF296; ORF297; ORF299; ORF300; ORF308; ORF314; ORF317; ORF318; ORF324; ORF342; ORF343; ORF355; ORF360; ORF374; ORF376; ORF386; ORF388; ORF392; ORF394; ORF395; ORF402; ORF405; ORF411; ORF415; ORF416;

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ORF422; ORF423; ORF429; ORF432; ORF441; ORF442; ORF444; ORF449; ORF452; ORF456; ORF460; ORF461; ORF465; ORF471; ORF472; ORF482; ORF489; ORF492; ORF494; ORF495; ORF502; ORF505; ORF506; ORF509; ORF516; ORF517; ORF520; ORF525; ORF533; ORF539; ORF549; ORF554; ORF557; ORF563; ORF570; ORF573; ORF581; ORF590; ORF591; ORF600; ORF607; ORF612; ORF613; ORF620; ORF626; ORF629; ORF630; ORF639; ORF644; ORF647; ORF656; ORF659; ORF661; ORF685; ORF687; ORF699; ORF700; ORF708; ORF716; ORF719; ORF725; ORF747; ORF749; ORF756; ORF765; ORF767; ORF794; ORF796; ORF797; ORF799; ORF801; ORF807; ORF821; ORF823; ORF826; ORF847; ORF853; ORF861; ORF870; ORF871; ORF875; ORF882; ORF888; ORF889; ORF898; ORF902; ORF903; ORF911; ORF916; ORF931; ORF939; ORF975; ORF976; ORF978; ORF983; ORF986; ORF987; ORF992; ORF993; ORF1000; ORF1002; ORF1008; ORF1019; ORF1022; ORF1032; ORF1034; ORF1046; ORF1054; ORF1060; ORF1071 and one of their representative fragments.
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Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF4; ORF6; ORF13; ORF20; ORF51; ORF71; ORF88; ORF118; ORF128; ORF132; ORF133; ORF158; ORF159; ORF174; ORF180; ORF189; ORF210; ORF211; ORF214; ORF215; ORF226; ORF229; ORF233; ORF235; ORF240; ORF246; ORF251; ORF255; ORF273; ORF354; ORF364; ORF369; ORF371; ORF397; ORF401; ORF409; ORF412; ORF419; ORF439; ORF453; ORF462; ORF490; ORF510; ORF511; ORF518; ORF535; ORF548; ORF550; ORF564; ORF565; ORF578; ORF579; ORF614; ORF631; ORF636; ORF650; ORF662; ORF667; ORF679; ORF681; ORF702; ORF727; ORF741; ORF763; ORF791; ORF792; ORF815; ORF816; ORF832; ORF846; ORF858; ORF865; ORF867; ORF868; ORF877; ORF891; ORF896; ORF907; ORF908; ORF918; ORF919; ORF932; ORF959; ORF977; ORF994; ORF998; ORF1024; ORF1028; ORF1042; ORF1067; ORF1070; ORF1073 and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* surface exposed polypeptide (e.g., an outer membrane protein) or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 2, ORF 3, ORF 21, ORF 22, ORF 23, ORF 53, ORF 77, ORF 187, ORF 203, ORF 383, ORF 477, ORF 478, ORF 479, ORF 481, ORF 482, ORF 483, ORF 484, ORF 485, ORF 486, ORF 487, ORF 488, ORF 489, ORF 490, ORF 571, ORF 572, ORF 573, ORF 593, ORF 670, ORF 693, ORF 742, ORF 749, ORF 801, ORF 817, ORF 818, ORF 819, ORF 820, ORF 851, ORF 902, ORF 923, ORF 1035, ORF 1036, ORF 1037, ORF 1038, ORF 1069, ORF 1070, ORF 1071, ORF 1073, ORF

1076, ORF 1095, ORF 1096, ORF 1141, ORF 1181, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* lipoprotein of one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences:

ORF 29, ORF 42, ORF 66, ORF 72, ORF 76, ORF 78, ORF 148, ORF 154, ORF 180, ORF 182, ORF 184, ORF 187, ORF 200, ORF 242, ORF 245, ORF 250, ORF 253, ORF 272, ORF 274, ORF 275, ORF 308, ORF 350, ORF 362, ORF 383, ORF 394, ORF 396, ORF 399,

ORF 422, ORF 488, ORF 535, ORF 568, ORF 573, ORF 578, ORF 593, ORF 607, ORF 625, ORF 662, ORF 669, ORF 688, ORF 690, ORF 716, ORF 773, ORF 778, ORF 781, ORF 783, ORF 788, ORF 817, ORF 848, ORF 851, ORF 853, ORF 857, ORF 875, ORF 877, ORF 886, ORF 898, ORF 902, ORF 923, ORF 938, ORF 976, ORF 978, ORF 990, ORF 1005, ORF 1021, ORF 1035, ORF 1069, ORF 1083, ORF 1088, ORF 1089, ORF 1091, ORF 1092, ORF 1095, ORF 1096, ORF 1100, ORF 1105, ORF 1108, ORF 1117, ORF 1120, ORF 1121, ORF 1124, ORF 1128, ORF 1133, ORF 1135, ORF 1139, ORF 1140, ORF 1157, ORF 1159, ORF 1163, ORF 1165, ORF 1167, ORF 1168, ORF 1169, ORF 1171, ORF 1173, ORF 1174, ORF 1177, ORF 1180, ORF 1181, ORF 1186, ORF 1194, ORF 1197, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide involved in lipopolysaccharide (LPS) biosynthesis, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 17, ORF 201, ORF 691, ORF 807, ORF 936, ORF 983, ORF 1019, ORF 1077 and one of their representative fragments.

Preferably the invention relates to additional LPS-related nucleotide sequences according to the invention, characterized in that they encode:

- 25 (a) a Chlamydia trachomatis KDO (3-deoxy-D-manno-octulosonic acid)-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 41, ORF 242, ORF 269, ORF 772, and one of their representative fragments;
- (b) a Chlamydia trachomatis phosphomannomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequence: ORF 139, and one of its representative fragments;
  - (c) a Chlamydia trachomatis phosphoglucomutase-related polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequence: ORF 567, and one of its representative fragments; and
- 35 (d) a Chlamydia trachomatis lipid A component-related polypeptide or one of its

representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 4, ORF 933, ORF 934, ORF 935, ORF 1185, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* Type III or other, non-Type III secreted polypeptides or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 180, ORF 181, ORF 207, ORF 208, ORF 372, ORF 391, ORF 399, ORF 477, ORF 486, ORF 749, ORF 758, ORF 819, ORF 878, ORF 888, ORF 896, ORF 897, ORF 900, ORF 902, ORF 923, ORF 1015, ORF 1018, ORF 1059, ORF 1060, ORF 1069, ORF 1071, ORF 1073, ORF 1076, ORF 1189, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide containing RGD (Arg-Gly-Asp) attachment sites or one of its representative fragments:

- 15 (a) RGD-containing proteins that are outer membrane proteins, are more likely to play a role in cell attachment. ORFs that encoded a protein containing an RGD sequence and also were classified as outer membrane proteins are ORF 488, ORF 489, ORF 571, ORF 572, ORF 573 or ORF 716, and its representative fragments.
- 20 (b) The outer membrane of Chlamydia is made of cysteine-rich proteins that form a network of both intra and inter molecular disulfide links. This contributes to the integrity of the membrane since Chlamydia lacks the peptidoglycan layer that other gram-negative bacteria have. Cysteine-rich proteins that have the RGD sequence are also considered to be potential vaccine candidates. Cysteine-rich proteins were defined as proteins that had more than 3.0% cysteine in their primary amino acid sequence, above the mean genomic ORF cysteine content. The corresponding ORF is: ORF 1144 and one of its representative fragments.
- (c) The outer membrane of Chlamydia may also contain small proteins that have cysteines in their N- and C-terminus that may contribute to the network formed by disulfide linkages.

  These proteins may be anchored in the outer membrane via their N-terminus and may have their C-terminus exposed, which then can interact with the host cells. Alternatively, these proteins may be anchored in the outer membrane via both N-and C-terminus and may have regions in the middle that may be exposed which can in turn interact with the host cells. ORFs encoding polypeptides that contain cysteines in their first 30 amino acids and also contain an RGD sequence are: ORF 101, ORF 122, ORF 308, ORF 488, ORF 489, ORF 571, ORF 572, ORF 573, ORF 651, ORF 679, ORF 680,

ORF 705, ORF 716, ORF 763, ORF 870, ORF 878, ORF 879, ORF 995, ORF 1028, ORF 1029, ORF 1176, and one of their representative fragments.

(d) RGD-containing ORFs homologous to RGD-containing ORFs from
 5 Chlamydia pneumoniae are:
 ORF 28, ORF 101, ORF 125, ORF 155, ORF 156, ORF 286, ORF 571, ORF 572, ORF 573, ORF 763, ORF 870, and one of their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* cell wall anchored surface polypeptide or one of its representative fragments, said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 662, ORF 681, ORF 1182, ORF 1192, and their representative fragments.

Preferably, the invention relates to the nucleotide sequences according to the 15 invention, characterized in that they encode Chlamydia trachomatis polypeptides not found in Chlamydia pneumoniae (Blastp P>e-10), said nucleotide sequences comprising a nucleotide sequence chosen from the following sequences: ORF 2, ORF 18, ORF 60, ORF 66, ORF 67, ORF 68, ORF 69, ORF 70, ORF 81, ORF 89, ORF 107, ORF 108, ORF 109, ORF 134, ORF 147, ORF 191, ORF 194, ORF 216, ORF 217, ORF 218, ORF 219, ORF 220, ORF 221, ORF 222, ORF 223, ORF 224, ORF 20 225, ORF 228, ORF 235, ORF 257, ORF 276, ORF 277, ORF 278, ORF 279, ORF 280, ORF 281, ORF 282, ORF 283, ORF 284, ORF 285, ORF 289, ORF 291, ORF 298, ORF 313, ORF 314, ORF 315, ORF 316, ORF 334, ORF 335, ORF 336, ORF 337, ORF 338, ORF 339, ORF 340, ORF 381, ORF 393, ORF 413, ORF 418, ORF 419, ORF 420, ORF 421, ORF 422, ORF 423, ORF 436, ORF 460, ORF 475, ORF 476, ORF 480, ORF 485, ORF 487, ORF 491, ORF 492, ORF 493, ORF 494, 25 ORF 496, ORF 500, ORF 504, ORF 514, ORF 527, ORF 559, ORF 569, ORF 570, ORF 575, ORF 580, ORF 582, ORF 593, ORF 598, ORF 632, ORF 640, ORF 651, ORF 671, ORF 690, ORF 694, ORF 698, ORF 710, ORF 722, ORF 723, ORF 724, ORF 770, ORF 771, ORF 782, ORF 783, ORF 784, ORF 790, ORF 795, ORF 798, ORF 805, ORF 810, ORF 817, ORF 829, ORF 830, ORF 864, ORF 866, ORF 876, ORF 887, ORF 892, ORF 899, ORF 913, ORF 921, ORF 933, ORF 938, ORF 30 949, ORF 956, ORF 1010, ORF 1017, ORF 1018, ORF 1027, ORF 1030, ORF 1037, ORF 1038, ORF 1047, ORF 1072, ORF 1074, ORF 1075, ORF 1078, ORF 1079, ORF 1081, ORF 1083, ORF 1084, ORF 1087, ORF 1088, ORF 1089, ORF 1091, ORF 1092, ORF 1094, ORF 1095, ORF 1096, ORF 1098, ORF 1104, ORF 1105, ORF 1106, ORF 1108, ORF 1110, ORF 1114, ORF 1115, ORF 1116, ORF 1117, ORF 1119, ORF 1128, ORF 1132, ORF 1133, ORF 1135, ORF 1136, ORF 1139, ORF 1140, ORF 1141, ORF 1142, ORF 1144, ORF 1148, ORF 1151, ORF 1155, ORF 1157, ORF 1159,

ORF 1161, ORF 1162, ORF 1165, ORF 1166, ORF 1167, ORF 1168, ORF 1169, ORF 1171, ORF 1172, ORF 1173, ORF 1174, ORF 1175, ORF 1176, ORF 1177, ORF 1178, ORF 1180, ORF 1181, ORF 1183, ORF 1184, ORF 1186, ORF 1187, ORF 1188, ORF 1192, ORF 1194, ORF 1197, and their representative fragments.

5 Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a Chlamydia trachomatis polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, such as for example triose phosphate isomerase or pyruvate kinase, and in that they comprise a nucleotide sequence chosen from the following sequences: 10 ORF10; ORF44; ORF45; ORF46; ORF47; ORF93; ORF101; ORF102; ORF103; ORF106; ORF107; ORF120; ORF121; ORF130; ORF135; ORF140; ORF143; ORF144; ORF145; ORF158; ORF159; ORF160; ORF161; ORF192; ORF193; ORF196; ORF197; ORF198; ORF199; ORF227; ORF229; ORF236; ORF236; ORF239; ORF243; ORF245; ORF264; ORF265; ORF297; ORF331; ORF333; ORF359; ORF360; ORF374; ORF404; ORF405; ORF410; ORF415; ORF415; ORF416; 15 ORF417; ORF432; ORF460; ORF461; ORF462; ORF495; ORF513; ORF515; ORF566; ORF566; ORF566; ORF689; ORF613; ORF645; ORF646; ORF647; ORF652; ORF653; ORF654; ORF672; ORF673; ORF674; ORF682; ORF684; ORF692; ORF700; ORF725; ORF801; ORF802; ORF835; ORF836; ORF867; ORF860; ORF861; ORF862; ORF863; ORF869; ORF925; ORF964; ORF983 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, such as for example CTP synthetase or GMP synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF142; ORF142; ORF169; ORF256; ORF268; ORF325; ORF352; ORF366; ORF435; ORF444; ORF528; ORF529; ORF530; ORF548; ORF549; ORF601; ORF602; ORF617; ORF619; ORF644; ORF745; ORF971; ORF972; ORF1023 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, such as for example DNA polymerases or DNA topoisomerases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF5; ORF12; ORF82; ORF96; ORF97; ORF98; ORF99; ORF100; ORF105; ORF118; ORF136; ORF137; ORF163; ORF190; ORF204; ORF259; ORF260; ORF262; ORF290; ORF300; ORF301; ORF302; ORF387; ORF427; ORF434; ORF441; ORF444; ORF471; ORF595; ORF596; ORF597;

ORF599; ORF600; ORF605; ORF612; ORF624; ORF625; ORF650; ORF657; ORF658; ORF702; ORF703; ORF704; ORF708; ORF719; ORF766; ORF767; ORF775; ORF779; ORF787; ORF788; ORF794; ORF841; ORF842; ORF883; ORF884; ORF907; ORF918; ORF924; ORF928; ORF929; ORF962; ORF962; ORF963; ORF969; ORF970; ORF975; ORF979; ORF995; ORF1031; ORF1032 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, such as for example serine hydroxymethyl transferase or the proteins which load amino acids onto transfer RNAs, and in that they comprise a nucleotide sequence chosen from the following sequences: ORF27; ORF41; ORF55; ORF56; ORF57; ORF59; ORF62; ORF63; ORF64; ORF65; ORF119; ORF132; ORF240; ORF241; ORF277; ORF278; ORF279; ORF382; ORF406; ORF428; ORF442; ORF446; ORF447; ORF453; ORF454; ORF541; ORF542; ORF591; ORF608; ORF609; ORF610; ORF618; ORF648; ORF649; ORF660; ORF661; ORF677; ORF717; ORF765; ORF797; ORF871; ORF875; ORF920; ORF922; ORF937; ORF998; ORF1020; ORF1021; ORF1034; ORF1044;

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, such as for example protein kinases or proteases, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF1046; ORF1049 and one of their representative fragments.

ORF21; ORF22; ORF23; ORF24; ORF25; ORF26; ORF75; ORF84; ORF86; ORF92; ORF133;
ORF151; ORF152; ORF157; ORF179; ORF209; ORF307; ORF326; ORF343; ORF344; ORF345;
ORF371; ORF429; ORF519; ORF557; ORF586; ORF587; ORF630; ORF656; ORF706; ORF707;
ORF730; ORF751; ORF752; ORF786; ORF847; ORF885; ORF923; ORF978; ORF1039; ORF1048 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, such as for example succinyl-CoA-synthesizing proteins or phosphatidylserine synthetase, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF4; ORF15; ORF16; ORF141; ORF173; ORF205; ORF205; ORF206; ORF207; ORF208; ORF312; ORF355; ORF415; ORF550; ORF558; ORF560; ORF561; ORF574; ORF577; ORF578; ORF590; ORF614; ORF772; ORF808; ORF809; ORF904; ORF905; ORF905; ORF933; ORF934; ORF934; ORF936 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or on of its representative fragments which is involved in the synthesis of the wall, such as for example KDO transferase, and the proteins responsible for the attachment of certain sugars onto the exposed proteins, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF87; ORF196; ORF242; ORF269; ORF628; ORF629; ORF634; ORF635; ORF637; ORF638; ORF1019 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, such as for example initiation factors, RNA polymerases or certain chaperone proteins, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF112; ORF113; ORF332; ORF212; ORF213; ORF350; ORF362; ORF363; ORF364; ORF407; ORF451; ORF546; ORF643; ORF744; ORF746; ORF833; ORF868; ORF981; ORF982; ORF1003; ORF1011; ORF1042 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* ribosomal polypeptide or one of its representative fragments, such as for example the ribosomal proteins L21, L27 and S10, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF114; ORF115; ORF116; ORF328; ORF361; ORF375; ORF445; ORF543; ORF584; ORF585; ORF743; ORF813; ORF941; ORF942; ORF944; ORF946; ORF947; ORF948; ORF950; ORF951; ORF952; ORF953; ORF954; ORF955; ORF955; ORF957; ORF958; ORF960; ORF961; ORF1040; ORF1041; ORF1043; ORF1063; ORF1064 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* transport polypeptide or one of its representative fragments, such as for example the proteins for transporting amino acids, sugars and certain oligopeptides, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF6; ORF50; ORF51; ORF80; ORF125; ORF126; ORF128; ORF129; ORF215; ORF246; ORF248; ORF249; ORF251; ORF252; ORF253; ORF255; ORF271; ORF275; ORF293; ORF309; ORF323; ORF324; ORF398; ORF401; ORF449; ORF511; ORF512; ORF564; ORF565; ORF667; ORF679; ORF680; ORF711; ORF712; ORF713; ORF714; ORF715; ORF730; ORF731; ORF736; ORF737; ORF738; ORF870; ORF908; ORF919; ORF977; ORF987; ORF988; ORF992; ORF993; ORF994; ORF1028; ORF1029 and one of their representative fragments.

invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the virulence process, such as for example the proteins analogous to the *Escherichia coli* vacB protein, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF20; ORF815; ORF816; ORF898; ORF1059; ORF1060 and one of their representative fragments.

Preferably, the invention also relates to the nucleotide sequences according to the invention, characterized in that they encode a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the secretory system and/or which is secreted, such as for example proteins homologous to proteins in the secretory system of certain bacteria such as the Salmonellae or the Yersiniae, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF758; ORF888; ORF889; ORF890; ORF891; ORF896; ORF897; ORF898 and one of their representative fragments.

Preferably, the invention also relates to nucleotide sequences according to the invention, characterized in that they encode a polypeptide specific to Chlamydiae or one of its representative fragments, and in that they comprise a nucleotide sequence chosen from the following sequences:

ORF22; ORF29; ORF31; ORF32; ORF34; ORF35; ORF39; ORF40; ORF43; ORF48; ORF49; ORF50; ORF52; ORF53; ORF54; ORF72; ORF77; ORF78; ORF87; ORF90; ORF95; ORF108; 20 ORF110; ORF121; ORF122; ORF123; ORF124; ORF127; ORF138; ORF144; ORF146; ORF153; ORF155; ORF164; ORF166; ORF175; ORF182; ORF184; ORF186; ORF187; ORF188; ORF202; ORF210; ORF247; ORF258; ORF266; ORF267; ORF270; ORF273; ORF274; ORF295; ORF296; ORF305; ORF306; ORF309; ORF318; ORF319; ORF322; ORF326; ORF342; ORF357; ORF376; ORF379; ORF380; ORF388; ORF390; ORF400; ORF431; ORF433; ORF438; ORF443; ORF456; 25 ORF457; ORF458; ORF464; ORF468; ORF470; ORF473; ORF486; ORF489; ORF497; ORF501; ORF503; ORF504; ORF508; ORF512; ORF521; ORF522; ORF523; ORF524; ORF533; ORF535; ORF536; ORF537; ORF538; ORF539; ORF540; ORF554; ORF563; ORF572; ORF579; ORF595; ORF603; ORF604; ORF606; ORF607; ORF615; ORF616; ORF622; ORF641; ORF642; ORF659; ORF668; ORF670; ORF693; ORF695; ORF696; ORF699; ORF703; ORF704; ORF716; ORF726; 30 ORF728; ORF739; ORF742; ORF747; ORF750; ORF751; ORF755; ORF757; ORF759; ORF761; ORF762; ORF763; ORF764; ORF773; ORF780; ORF781; ORF789; ORF800; ORF803; ORF804; ORF818; ORF820; ORF822; ORF823; ORF824; ORF827; ORF828; ORF839; ORF849; ORF850; ORF851; ORF852; ORF855; ORF856; ORF857; ORF858; ORF859; ORF860; ORF861; ORF862; ORF863; ORF865; ORF869; ORF870; ORF871; ORF872; ORF873; ORF874; ORF875; 35 ORF877; ORF878; ORF880; ORF882; ORF884; ORF886; ORF893; ORF901; ORF906; ORF910;

ORF912; ORF915; ORF916; ORF917; ORF926; ORF929; ORF933; ORF965; ORF967; ORF968; ORF984; ORF986; ORF989; ORF990; ORF996; ORF997; ORF1001; ORF1002; ORF1013; ORF1016; ORF1031; ORF1033; ORF1035; ORF1049; ORF1051; ORF1052; ORF1054; ORF1056; ORF1057; ORF1058; ORF1062; ORF1070; ORF1071; ORF1073 and one of their representative fragments.

Also forming part of the invention are polypeptides encoded by the polynucleotides of the invention, as well as fusion polypeptides comprising such polypeptides. In one embodiment, the polypeptides and fusion polypeptides immunoreact with seropositive serum of an individual infected with *Chlamydia trachomatis*. For example, described below, are polypeptide sequences exhibiting particularly preferable characteristics. For each group of preferred polypeptides described below, it is to be understood that in addition to the individual polypeptides listed, in instances wherein such polypeptides are encoded as part of «combined» ORFs, such «combined» polypeptides are also to be included within the preferred group.

The subject of the invention is also a polypeptide according to the invention, characterized in that it is a polypeptide of the cellular envelope, preferably of the outer cellular envelope, of *Chlamydia trachomatis* or one of its representative fragments. According to the invention, the said polypeptide is preferably chosen from the polypeptides having the following sequences:

SEQ ID No. 3; SEQ ID No. 19; SEQ ID No. 51; SEQ ID No. 189; SEQ ID No. 212; SEQ ID No. 213; 20 SEQ ID No. 324; SEQ ID No. 477; SEQ ID No. 478; SEQ ID No. 479; SEQ ID No. 481; SEQ ID No. 482; SEQ ID No. 483; SEQ ID No. 484; SEQ ID No. 486; SEQ ID No. 488; SEQ ID No. 489; SEQ ID No. 490; SEQ ID No. 572; SEQ ID No. 573; SEQ ID No. 742; SEQ ID No. 817; SEQ ID No. 818; SEQ ID No. 820; SEQ ID No. 1035; SEQ ID No. 1036; SEQ ID No. 1037; SEQ ID No. 1038; SEQ ID No. 1070; SEQ ID No. 1071; SEQ ID No. 1073 and 25 one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 1 and 3 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 2; SEQ ID No. 3; SEQ ID No. 5; SEQ ID No. 8; SEQ ID No. 9; SEQ ID No. 10;
SEQ ID No. 11; SEQ ID No. 12; SEQ ID No. 17; SEQ ID No. 21; SEQ ID No. 26; SEQ ID No. 27;
SEQ ID No. 28; SEQ ID No. 29; SEQ ID No. 30; SEQ ID No. 31; SEQ ID No. 33; SEQ ID No. 35;
SEQ ID No. 37; SEQ ID No. 39; SEQ ID No. 40; SEQ ID No. 41; SEQ ID No. 42; SEQ ID No. 43;
SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 46; SEQ ID No. 47; SEQ ID No. 48; SEQ ID No. 49;
SEQ ID No. 52; SEQ ID No. 53; SEQ ID No. 55; SEQ ID No. 56; SEQ ID No. 58; SEQ ID No. 65;

SEQ ID No. 66; SEQ ID No. 68; SEQ ID No. 70; SEQ ID No. 74; SEQ ID No. 75; SEQ ID No. 76; SEQ ID No. 78; SEQ ID No. 79; SEQ ID No. 81; SEQ ID No. 82; SEQ ID No. 83; SEQ ID No. 86; SEQ ID No. 91; SEQ ID No. 92; SEQ ID No. 94; SEQ ID No. 97; SEQ ID No. 100; SEQ ID No. 102; SEQ ID No. 103; SEQ ID No. 105; SEQ ID No. 106; SEQ ID No. 107; SEQ ID No. 109; 5 SEQ ID No. 110; SEQ ID No. 111; SEQ ID No. 112; SEQ ID No. 113; SEQ ID No. 114; SEQ ID No. 115; SEQ ID No. 116; SEQ ID No. 117; SEQ ID No. 120; SEQ ID No. 122; SEQ ID No. 123; SEQ ID No. 130; SEQ ID No. 134; SEQ ID No. 135; SEQ ID No. 137; SEQ ID No. 140; SEQ ID No. 141; SEQ ID No. 143; SEQ ID No. 144; SEQ ID No. 145; SEQ ID No. 147; SEQ ID No. 148; SEQ ID No. 149; SEQ ID No. 150; SEQ ID No. 151; 10 SEQ ID No. 155; SEQ ID No. 156; SEQ ID No. 162; SEQ ID No. 163; SEQ ID No. 164; SEQ ID No. 165; SEQ ID No. 166; SEQ ID No. 167; SEQ ID No. 168; SEQ ID No. 169; SEQ ID No. 170; SEQ ID No. 171; SEQ ID No. 173; SEQ ID No. 175; SEQ ID No. 176; SEQ ID No. 177; SEQ ID No. 181; SEQ ID No. 183; SEQ ID No. 184; SEQ ID No. 186; SEQ ID No. 187; SEQ ID No. 188; SEQ ID No. 190; SEQ ID No. 191; SEQ ID No. 192; SEQ ID No. 194; SEQ ID No. 195; SEQ ID No. 196; SEQ ID No. 197; SEQ ID No. 198; SEQ ID No. 199; SEQ ID No. 201; SEQ ID No. 202; SEQ ID No. 204; SEQ ID No. 206; SEQ ID No. 207; SEQ ID No. 209; SEQ ID No. 212; SEQ ID No. 213; SEQ ID No. 217; SEQ ID No. 219; SEQ ID No. 220; SEQ ID No. 221; SEQ ID No. 222; SEQ ID No. 223; SEQ ID No. 224; SEQ ID No. 225; SEQ ID No. 227; SEQ ID No. 228; SEQ ID No. 231; SEQ ID No. 232; SEQ ID No. 234; SEQ ID No. 236; SEQ ID No. 237; SEQ ID No. 243; SEQ ID No. 244; SEQ ID No. 245; SEQ ID No. 247; SEQ ID No. 248; SEQ ID No. 249; SEQ ID No. 252; SEQ ID No. 254; SEQ ID No. 257; SEQ ID No. 260; SEQ ID No. 261; SEQ ID No. 263; SEQ ID No. 265; SEQ ID No. 266; SEQ ID No. 267; SEQ ID No. 270; SEQ ID No. 271; SEQ ID No. 272; SEQ ID No. 274; SEQ ID No. 276; SEQ ID No. 277; 25 SEQ ID No. 278; SEQ ID No. 279; SEQ ID No. 282; SEQ ID No. 283; SEQ ID No. 284; SEQ ID No. 285; SEQ ID No. 287; SEQ ID No. 289; SEQ ID No. 290; SEQ ID No. 291; SEQ ID No. 294; SEQ ID No. 298; SEQ ID No. 305; SEQ ID No. 306; SEQ ID No. 310; SEQ ID No. 311; SEQ ID No. 313; SEQ ID No. 315; SEQ ID No. 316; SEQ ID No. 319; SEQ ID No. 320; SEQ ID No. 322; SEQ ID No. 323; SEQ ID No. 325; SEQ ID No. 326; 30 SEQ ID No. 327; SEQ ID No. 328; SEQ ID No. 330; SEQ ID No. 331; SEQ ID No. 332; SEQ ID No. 333; SEQ ID No. 334; SEQ ID No. 335; SEQ ID No. 336; SEQ ID No. 338; SEQ ID No. 339; SEQ ID No. 340; SEQ ID No. 341; SEQ ID No. 344; SEQ ID No. 345; SEQ ID No. 348; SEQ ID No. 349; SEQ ID No. 350; SEQ ID No. 351; SEQ ID No. 352; SEQ ID No. 353; SEQ ID No. 356; SEQ ID No. 357; SEQ ID No. 358; SEQ ID No. 361; SEQ ID No. 362; SEQ ID No. 366; SEQ ID No. 367; SEQ ID No. 368; SEQ ID No. 370;

	SEQ ID No. 372;	SEQ ID No. 373;	SEQ ID No. 375;	SEQ ID No. 377;	SEQ ID No. 378;
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	SEQ ID No. 385;	SEQ ID No. 387;	SEQ ID No. 389;	SEQ ID No. 390;	SEQ ID No. 391;
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5	SEQ ID No. 404;	SEQ ID No. 406;	SEQ ID No. 407;	SEQ ID No. 413;	SEQ ID No. 414;
	SEQ ID No. 417;	SEQ ID No. 418;	SEQ ID No. 420;	SEQ ID No. 421;	SEQ ID No. 424;
	SEQ ID No. 426;	SEQ ID No. 427;	SEQ ID No. 428;	SEQ ID No. 430;	SEQ ID No. 433;
	SEQ ID No. 434;	SEQ ID No. 435;	SEQ ID No. 436;	SEQ ID No. 437;	SEQ ID No. 440;
	SEQ ID No. 443;	SEQ ID No. 446;	SEQ ID No. 448;	SEQ ID No. 450;	SEQ ID No. 451;
10	SEQ ID No. 454;	SEQ ID No. 455;	SEQ ID No. 457;	SEQ ID No. 458;	SEQ ID No. 459;
	SEQ ID No. 463;	SEQ ID No. 464;	SEQ ID No. 466;	SEQ ID No. 467;	SEQ ID No. 468;
	SEQ ID No. 469;	SEQ ID No. 470;	SEQ ID No. 473;	SEQ ID No. 474;	SEQ ID No. 475;
	SEQ ID No. 476;	SEQ ID No. 477;	SEQ ID No. 479;	SEQ ID No. 480;	SEQ ID No. 481;
	SEQ ID No. 483;	SEQ ID No. 484;	SEQ ID No. 485;	SEQ ID No. 486;	SEQ ID No. 487;
15	SEQ ID No. 488;	SEQ ID No. 491;	SEQ ID No. 493;	SEQ ID No. 496;	SEQ ID No. 497;
	SEQ ID No. 498;	SEQ ID No. 500;	SEQ ID No. 501;	SEQ ID No. 503;	SEQ ID No. 504;
	SEQ ID No. 508;	SEQ ID No. 512;	SEQ ID No. 513;	SEQ ID No. 514;	SEQ ID No. 519;
	SEQ ID No. 521;	SEQ ID No. 523;	SEQ ID No. 524;	SEQ ID No. 526;	SEQ ID No. 527;
	SEQ ID No. 529;	SEQ ID No. 530;	SEQ ID No. 531;	SEQ ID No. 532;	SEQ ID No. 534;
20	SEQ ID No. 536;	SEQ ID No. 537;	SEQ ID No. 538;	SEQ ID No. 540;	SEQ ID No. 541;
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	SEQ ID No. 547;	SEQ ID No. 551;	SEQ ID No. 552;	SEQ ID No. 553;	SEQ ID No. 555;
	SEQ ID No. 558;	SEQ ID No. 559;	SEQ ID No. 560;	SEQ ID No. 561;	SEQ ID No. 562;
	SEQ ID No. 566;	SEQ ID No. 567;	SEQ ID No. 568;	SEQ ID No. 569;	SEQ ID No. 571;
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	SEQ ID No. 593;	SEQ ID No. 595;	SEQ ID No. 596;	SEQ ID No. 597;	SEQ ID No. 599;
	SEQ ID No. 601;	SEQ ID No. 602;	SEQ ID No. 603;	SEQ ID No. 604;	SEQ ID No. 608;
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30	SEQ ID No. 617;	SEQ ID No. 618;	SEQ ID No. 621;	SEQ ID No. 622;	SEQ ID No. 623;
	SEQ ID No. 624;	SEQ ID No. 625;	SEQ ID No. 628;	SEQ ID No. 632;	SEQ ID No. 633;
	SEQ ID No. 634;	SEQ ID No. 635;	SEQ ID No. 637;	SEQ ID No. 638;	SEQ ID No. 640;
	SEQ ID No. 641;	SEQ ID No. 643;	SEQ ID No. 646;	SEQ ID No. 648;	SEQ ID No. 649;
	SEQ ID No. 651;	SEQ ID No. 652;	SEQ ID No. 653;	SEQ ID No. 654;	SEQ ID No. 655;
35	SEQ ID No. 658;	SEQ ID No. 664;	SEQ ID No. 665;	SEQ ID No. 666;	SEQ ID No. 668;

	SEQ ID No. 669;	SEQ ID No. 670;	SEQ ID No. 671;	SEQ ID No. 672;	SEQ ID No. 673;
	SEQ ID No. 674;	SEQ ID No. 676;	SEQ ID No. 677;	SEQ ID No. 678;	SEQ ID No. 680;
	SEQ ID No. 682;	SEQ ID No. 683;	SEQ ID No. 684;	SEQ ID No. 686;	SEQ ID No. 688;
	SEQ ID No. 689;	SEQ ID No. 690;	SEQ ID No. 691;	SEQ ID No. 692;	SEQ ID No. 693;
5	SEQ ID No. 695;	SEQ ID No. 696;	SEQ ID No. 698;	SEQ ID No. 701;	SEQ ID No. 703;
	SEQ ID No. 704;	SEQ ID No. 705;	SEQ ID No. 706;	SEQ ID No. 707;	SEQ ID No. 709;
	SEQ ID No. 710;	SEQ ID No. 711;	SEQ ID No. 712;	SEQ ID No. 713;	SEQ ID No. 714;
	SEQ ID No. 715;	SEQ ID No. 717;	SEQ ID No. 718;	SEQ ID No. 720;	SEQ ID No. 721;
	SEQ ID No. 722;	SEQ ID No. 724;	SEQ ID No. 726;	SEQ ID No. 728;	SEQ ID No. 729;
10	SEQ ID No. 730;	SEQ ID No. 731;	SEQ ID No. 732;	SEQ ID No. 733;	SEQ ID No. 734;
	SEQ ID No. 737;	SEQ ID No. 738;	SEQ ID No. 739;	SEQ ID No. 740;	SEQ ID No. 742;
	SEQ ID No. 743;	SEQ ID No. 744;	SEQ ID No. 745;	SEQ ID No. 746;	SEQ ID No. 748;
	SEQ ID No. 750;	SEQ ID No. 751;	SEQ ID No. 752;	SEQ ID No. 753;	SEQ ID No. 754;
	SEQ ID No. 755;	SEQ ID No. 757;	SEQ ID No. 758;	SEQ ID No. 759;	SEQ ID No. 760;
15	SEQ ID No. 764;	SEQ ID No. 766;	SEQ ID No. 768;	SEQ ID No. 769;	SEQ ID No. 771;
	SEQ ID No. 772;	SEQ ID No. 773;	SEQ ID No. 774;	SEQ ID No. 775;	SEQ ID No. 776;
	SEQ ID No. 777;	SEQ ID No. 778;	SEQ ID No. 779;	SEQ ID No. 780;	SEQ ID No. 781;
	SEQ ID No. 782;	SEQ ID No. 783;	SEQ ID No. 786;	SEQ ID No. 787;	SEQ ID No. 788;
	SEQ ID No. 789;	SEQ ID No. 790;	SEQ ID No. 793;	SEQ ID No. 798;	SEQ ID No. 800;
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	SEQ ID No. 810;	SEQ ID No. 811;	SEQ ID No. 813;	SEQ ID No. 814;	SEQ ID No. 817;
	SEQ ID No. 820;	SEQ ID No. 822;	SEQ ID No. 824;	SEQ ID No. 825;	SEQ ID No. 827;
	SEQ ID No. 828;	SEQ ID No. 829;	SEQ ID No. 830;	SEQ ID No. 833;	SEQ ID No. 834;
	SEQ ID No. 835;	SEQ ID No. 837;	SEQ ID No. 838;	SEQ ID No. 839;	SEQ ID No. 840;
25	SEQ ID No. 841;	SEQ ID No. 842;	SEQ ID No. 843;	SEQ ID No. 845;	SEQ ID No. 848;
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	SEQ ID No. 872;	SEQ ID No. 873;	SEQ ID No. 874;	SEQ ID No. 878;	SEQ ID No. 879;
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	SEQ ID No. 886;	SEQ ID No. 887;	SEQ ID No. 892;	SEQ ID No. 893;	SEQ ID No. 894;
	SEQ ID No. 895;	SEQ ID No. 897;	SEQ ID No. 899;	SEQ ID No. 900;	SEQ ID No. 901;
	SEQ ID No. 904;	SEQ ID No. 906;	SEQ ID No. 909;	SEQ ID No. 910;	SEQ ID No. 912;
	SEQ ID No. 914;	SEQ ID No. 917;	SEQ ID No. 920;	SEQ ID No. 921;	SEQ ID No. 922;
35	SEQ ID No. 923;	SEQ ID No. 924;	SEQ ID No. 925;	SEQ ID No. 926;	SEQ ID No. 927;

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SEQ ID No. 930;
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    SEQ ID No. 937;
                       SEQ ID No. 940;
                                         SEQ ID No. 941;
                                                            SEQ ID No. 942;
                                                                              SEQ ID No. 943;
    SEQ ID No. 944;
                       SEQ ID No. 945;
                                         SEQ ID No. 947;
                                                            SEQ ID No. 948;
                                                                              SEQ ID No. 951;
    SEQ ID No. 952;
                       SEQ ID No. 953;
                                         SEQ ID No. 954;
                                                            SEQ ID No. 955;
                                                                              SEQ ID No. 956;
 5 SEQ ID No. 957;
                       SEQ ID No. 958;
                                         SEQ ID No. 960;
                                                            SEQ ID No. 961;
                                                                              SEQ ID No. 962;
    SEQ ID No. 963;
                       SEQ ID No. 964;
                                         SEQ ID No. 966;
                                                            SEQ ID No. 967;
                                                                              SEQ ID No. 969;
    SEQ ID No. 970;
                       SEQ ID No. 971;
                                         SEQ ID No. 973;
                                                           SEQ ID No. 974;
                                                                              SEQ ID No. 979;
                       SEQ ID No. 981;
    SEQ ID No. 980;
                                         SEQ ID No. 982;
                                                           SEQ ID No. 984;
                                                                              SEQ ID No. 988;
    SEQ ID No. 989;
                       SEQ ID No. 990;
                                         SEQ ID No. 991;
                                                           SEQ ID No. 995;
                                                                              SEQ ID No. 996;
10 SEQ ID No. 999;
                     SEQ ID No. 1001;
                                        SEQ ID No. 1003;
                                                                             SEQ ID No. 1005;
                                                          SEQ ID No. 1004;
    SEQ ID No. 1006;
                      SEQ ID No. 1007;
                                        SEQ ID No. 1009;
                                                          SEQ ID No. 1010;
                                                                            SEQ ID No. 1011;
    SEQ ID No. 1012;
                      SEQ ID No. 1013;
                                        SEQ ID No. 1014;
                                                          SEQ ID No. 1016;
                                                                            SEQ ID No. 1017;
    SEQ ID No. 1018;
                      SEQ ID No. 1020;
                                        SEQ ID No. 1021;
                                                          SEQ ID No. 1025;
                                                                            SEQ ID No. 1026;
                      SEQ ID No. 1029;
    SEQ ID No. 1027;
                                        SEQ ID No. 1030;
                                                          SEQ ID No. 1031;
                                                                            SEQ ID No. 1035;
   SEQ ID No. 1036;
                      SEQ ID No. 1037;
                                        SEQ ID No. 1038;
                                                          SEQ ID No. 1039;
                                                                            SEQ ID No. 1040;
    SEQ ID No. 1044;
                      SEQ ID No. 1045;
                                        SEQ ID No. 1047;
                                                          SEQ ID No. 1048;
                                                                            SEQ ID No. 1050;
    SEQ ID No. 1051;
                      SEQ ID No. 1052;
                                        SEQ ID No. 1053;
                                                          SEQ ID No. 1055;
                                                                            SEQ ID No. 1056;
    SEQ ID No. 1057;
                      SEQ ID No. 1058;
                                        SEQ ID No. 1061;
                                                          SEQ ID No. 1062; SEQ ID No. 1063;
    SEQ ID No. 1064; SEQ ID No. 1065; SEQ ID No. 1066; SEQ ID No. 1068; SEQ ID No. 1069;
   SEQ ID No. 1072; SEQ ID No. 1074; SEQ ID No. 1076 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having between 4 and 6 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 7; SEQ ID No. 14; SEQ ID No. 16; SEQ ID No. 32; SEQ ID No. 34; SEQ ID No. 36; SEQ ID No. 38; SEQ ID No. 50; SEQ ID No. 57; SEQ ID No. 59; SEQ ID No. 61; SEQ ID No. 62; SEQ ID No. 63; SEQ ID No. 64; SEQ ID No. 67; SEQ ID No. 69; SEQ ID No. 72; SEQ ID No. 77; SEQ ID No. 80; SEQ ID No. 84; SEQ ID No. 87; SEQ ID No. 93; SEQ ID No. 95; SEQ ID No. 99; SEQ ID No. 108; SEQ ID No. 119; SEQ ID No. 125; SEQ ID No. 129; SEQ ID No. 126; SEQ ID No. 131; SEQ ID No. 136; SEQ ID No. 139; SEQ ID No. 146; SEQ ID No. 152; SEQ ID No. 154; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 172; SEQ ID No. 179; SEQ ID No. 182; SEQ ID No. 185; SEQ ID No. 200; SEQ ID No. 203; SEQ ID No. 205; SEQ ID No. 239; SEQ ID No. 242; SEQ ID No. 250; SEQ ID No. 253; SEQ ID No. 256; SEQ ID No. 259; SEQ ID No. 262; SEQ ID No. 268; SEQ ID No. 275; SEQ ID No. 281; 35 SEQ ID No. 286; SEQ ID No. 288; SEQ ID No. 292; SEQ ID No. 295; SEQ ID No. 296;

	SEQ ID No. 297;	SEQ ID No. 299;	SEQ ID No. 300;	SEQ ID No. 308;	SEQ ID No. 314;
	SEQ ID No. 317;	SEQ ID No. 318;	SEQ ID No. 324;	SEQ ID No. 342;	SEQ ID No. 343;
	SEQ ID No. 355;	SEQ ID No. 360;	SEQ ID No. 374;	SEQ ID No. 376;	SEQ ID No. 386;
	SEQ ID No. 388;	SEQ ID No. 392;	SEQ ID No. 394;	SEQ ID No. 395;	SEQ ID No. 402;
5	SEQ ID No. 405;	SEQ ID No. 411;	SEQ ID No. 415;	SEQ ID No. 416;	SEQ ID No. 422;
	SEQ ID No. 423;	SEQ ID No. 429;	SEQ ID No. 432;	SEQ ID No. 441;	SEQ ID No. 442;
	SEQ ID No. 444;	SEQ ID No. 449;	SEQ ID No. 452;	SEQ ID No. 456;	SEQ ID No. 460;
	SEQ ID No. 461;	SEQ ID No. 465;	SEQ ID No. 471;	SEQ ID No. 472;	SEQ ID No. 482;
	SEQ ID No. 489;	SEQ ID No. 492;	SEQ ID No. 494;	SEQ ID No. 495;	SEQ ID No. 502;
10	SEQ ID No. 505;	SEQ ID No. 506;	SEQ ID No. 509;	SEQ ID No. 516;	SEQ ID No. 517;
	SEQ ID No. 520;	SEQ ID No. 525;	SEQ ID No. 533;	SEQ ID No. 539;	SEQ ID No. 549;
	SEQ ID No. 554;	SEQ ID No. 557;	SEQ ID No. 563;	SEQ ID No. 570;	SEQ ID No. 573;
	SEQ ID No. 581;	SEQ ID No. 590;	SEQ ID No. 591;	SEQ ID No. 600;	SEQ ID No. 607;
	SEQ ID No. 612;	SEQ ID No. 613;	SEQ ID No. 620;	SEQ ID No. 626;	SEQ ID No. 629;
15	SEQ ID No. 630;	SEQ ID No. 639;	SEQ ID No. 644;	SEQ ID No. 647;	SEQ ID No. 656;
	SEQ ID No. 659;	SEQ ID No. 661;	SEQ ID No. 685;	SEQ ID No. 687;	SEQ ID No. 699;
	SEQ ID No. 700;	SEQ ID No. 708;	SEQ ID No. 716;	SEQ ID No. 719;	SEQ ID No. 725;
	SEQ ID No. 747;	SEQ ID No. 749;	SEQ ID No. 756;	SEQ ID No. 765;	SEQ ID No. 767;
	SEQ ID No. 794;	SEQ ID No. 796;	SEQ ID No. 797;	SEQ ID No. 799;	SEQ ID No. 801;
20	SEQ ID No. 807;	SEQ ID No. 821;	SEQ ID No. 823;	SEQ ID No. 826;	SEQ ID No. 847;
	SEQ ID No. 853;	SEQ ID No. 861;	SEQ ID No. 870;	SEQ ID No. 871;	SEQ ID No. 875;
	SEQ ID No. 882;	SEQ ID No. 888;	SEQ ID No. 889;	SEQ ID No. 898;	SEQ ID No. 902;
	SEQ ID No. 903;	SEQ ID No. 911;	SEQ ID No. 916;	SEQ ID No. 931;	SEQ ID No. 939;
	SEQ ID No. 975;	SEQ ID No. 976;	SEQ ID No. 978;	SEQ ID No. 983;	SEQ ID No. 986;
25	SEQ ID No. 987;	SEQ ID No. 992;	SEQ ID No. 993;	SEQ ID No. 1000;	SEQ ID No. 1002;
	SEQ ID No. 1008;	SEQ ID No. 1019;	SEQ ID No. 1022;	SEQ ID No. 1032;	SEQ ID No. 1034;
	SEQ ID No. 1046;	SEQ ID No. 1054;	SEQ ID No. 1060;	SEQ ID No. 1071	and one of their
	representative fragr	ments.		•	
	D	familia di :	•		

Preferably, the invention relates to a polypeptide according to the invention,

characterized in that it is a *Chlamydia trachomatis* transmembrane polypeptide or one of its representative fragments, having at least 7 transmembrane domains, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 4; SEQ ID No. 6; SEQ ID No. 13; SEQ ID No. 20; SEQ ID No. 51; SEQ ID No. 71; SEQ ID No. 88; SEQ ID No. 118; SEQ ID No. 128; SEQ ID No. 132; SEQ ID No. 133; SEQ ID No. 158; SEQ ID No. 159; SEQ ID No. 174;

SEQ ID No. 180; SEQ ID No. 189; SEQ ID No. 210; SEQ ID No. 211; SEQ ID No. 214;

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SEQ ID No. 215;
                        SEQ ID No. 226;
                                          SEQ ID No. 229;
                                                            SEQ ID No. 233;
                                                                                SEQ ID No. 235;
     SEQ ID No. 240;
                        SEQ ID No. 246;
                                          SEQ ID No. 251;
                                                             SEQ ID No. 255;
                                                                               SEQ ID No. 273;
     SEQ ID No. 354;
                       SEQ ID No. 364;
                                          SEQ ID No. 369;
                                                            SEQ ID No. 371;
                                                                               SEQ ID No. 397;
     SEQ ID No. 401;
                       SEQ ID No. 409;
                                          SEQ ID No. 412;
                                                            SEQ ID No. 419;
                                                                               SEQ ID No. 439;
 5 SEQ ID No. 453;
                       SEQ ID No. 462;
                                          SEQ ID No. 490;
                                                            SEQ ID No. 510;
                                                                               SEQ ID No. 511;
    SEQ ID No. 518;
                       SEQ ID No. 535;
                                          SEQ ID No. 548;
                                                            SEQ ID No. 550;
                                                                               SEQ ID No. 564;
    SEQ ID No. 565;
                       SEQ ID No. 578;
                                          SEQ ID No. 579;
                                                            SEQ ID No. 614;
                                                                               SEQ ID No. 631;
    SEQ ID No. 636;
                       SEQ ID No. 650;
                                          SEQ ID No. 662;
                                                            SEQ ID No. 667;
                                                                               SEQ ID No. 679;
    SEQ ID No. 681;
                       SEQ ID No. 702;
                                          SEQ ID No. 727;
                                                            SEQ ID No. 741;
                                                                               SEQ ID No. 763;
10 SEQ ID No. 791;
                       SEQ ID No. 792;
                                          SEQ ID No. 815;
                                                            SEQ ID No. 816;
                                                                               SEQ ID No. 832;
    SEQ ID No. 846;
                       SEQ ID No. 858;
                                          SEQ ID No. 865;
                                                            SEQ ID No. 867;
                                                                               SEQ ID No. 868;
    SEQ ID No. 877;
                       SEQ ID No. 891;
                                         SEQ ID No. 896;
                                                            SEQ ID No. 907;
                                                                               SEQ ID No. 908;
                       SEQ ID No. 919;
    SEQ ID No. 918;
                                         SEQ ID No. 932;
                                                            SEQ ID No. 959;
                                                                               SEQ ID No. 977;
    SEQ ID No. 994;
                      SEQ ID No. 998;
                                        SEQ ID No. 1024; SEQ ID No. 1028;
                                                                             SEQ ID No. 1042;
15 SEQ ID No. 1067; SEQ ID No. 1070; SEQ ID No. 1073 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* surface exposed polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 2, SEQ ID No. 3, SEQ ID No. 21, SEQ ID No. 22, SEQ ID No. 23, SEQ ID No. 53, SEQ ID No. 77, SEQ ID No. 187, SEQ ID No. 203, SEQ ID No. 383, SEQ ID No. 477, SEQ ID No. 478, SEQ ID No. 479, SEQ ID No. 481, SEQ ID No. 482, SEQ ID No. 483, SEQ ID No. 484, SEQ ID No. 485, SEQ ID No. 486, SEQ ID No. 487, SEQ ID No. 488, SEQ ID No. 489, SEQ ID No. 490, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 593, SEQ ID No. 670, SEQ ID No. 693, SEQ ID No. 742, SEQ ID No. 749, SEQ ID No. 801, SEQ ID No. 817, SEQ ID No. 818, SEQ ID No. 819, SEQ ID No. 820, SEQ ID No. 851, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 1035, SEQ ID No. 1036, SEQ ID No. 1037, SEQ ID No. 1038, SEQ ID No. 1069, SEQ ID No. 1070, SEQ ID No. 1071, SEQ ID No. 1073, SEQ ID No. 1076, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1141, SEQ ID No. 1181, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* lipoprotein or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 29, SEQ ID No. 42, SEQ ID No. 66, SEQ ID No. 72, SEQ ID No. 76, SEQ ID No. 78, SEQ ID No. 148, SEQ ID No. 154, SEQ ID No. 180, SEQ ID No. 182, SEQ ID No. 184, SEQ ID No. 187, SEQ ID No. 200, SEQ ID No. 242, SEQ ID No. 245, SEQ ID No. 250, SEQ ID No. 253, SEQ ID No. 272, SEQ ID No. 274, SEQ ID No. 275, SEQ ID No. 308, SEQ ID No. 350, SEQ ID No. 362,

SEQ ID No. 383, SEQ ID No. 394, SEQ ID No. 396, SEQ ID No. 399, SEQ ID No. 422, SEQ ID No. 488, SEQ ID No. 535, SEQ ID No. 568, SEQ ID No. 573, SEQ ID No. 578, SEQ ID No. 593, SEQ ID No. 607, SEQ ID No. 625, SEQ ID No. 662, SEQ ID No. 669, SEQ ID No. 688, SEQ ID No. 690, SEQ ID No. 716, SEQ ID No. 773, SEQ ID No. 778, SEQ ID No. 781, SEQ ID No. 783, SEQ ID No. 788, SEQ ID No. 817, SEQ ID No. 848, SEQ ID No. 851, SEQ ID No. 853, SEQ ID No. 857, SEQ ID No. 875, SEQ ID No. 877, SEQ ID No. 886, SEQ ID No. 898, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 938, SEQ ID No. 976, SEQ ID No. 978, SEQ ID No. 990, SEQ ID No. 1005, SEQ ID No. 1021, SEQ ID No. 1035, SEQ ID No. 1069, SEQ ID No. 1083, SEQ ID No. 1088, SEQ ID No. 1089, SEQ ID No. 1091, SEQ ID No. 1092, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1100, SEQ ID No. 1105, SEQ ID No. 1108, SEQ ID No. 1117, SEQ ID No. 1120, SEQ ID No. 1121, SEQ ID No. 1124, SEQ ID No. 1128, SEQ ID No. 1133, SEQ ID No. 1135, SEQ ID No. 1139, SEQ ID No. 1140, SEQ ID No. 1157, SEQ ID No. 1169, SEQ ID No. 1163, SEQ ID No. 1165, SEQ ID No. 1167, SEQ ID No. 1169, SEQ ID No. 1171, SEQ ID No. 1173, SEQ ID No. 1174, SEQ ID No. 1177, SEQ ID No. 1180, SEQ ID No. 1181, SEQ ID No. 1186, SEQ ID No. 1194, SEQ ID No. 1197, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide involved in lipopolysaccharide (LPS) biosynthesis, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 17, SEQ ID No. 201, SEQ ID No. 691, SEQ ID No. 807, SEQ ID No. 936, SEQ ID No. 983, SEQ ID No. 1019, SEQ ID No. 1077, and their representative fragments.

Preferably, the invention relates to additional LPS-related polypeptides according to the invention, in that it is:

- (a) a Chlamydia trachomatis KDO (3-deoxy-D-manno-octylosonic acid)-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides
   25 having the following sequences: SEQ ID No. 41, SEQ ID No. 242, SEQ ID No. 269, SEQ ID No. 772, and one of their representative fragments;
  - (b) a Chlamydia trachomatis phosphomannomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequence: SEQ ID No. 139, and its representative fragments;
  - (c) a Chlamydia trachomatis phosphoglucomutase-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequence: SEQ ID No. 567 and its representative fragments; and
- (d) a Chlamydia trachomatis lipid A component-related polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following
   sequences: SEQ ID No. 4, SEQ ID No. 933, SEQ ID No. 934, SEQ ID No. 935, SEQ ID No. 1185,

35

and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments that contains an RGD sequence and is also an outer membrane protein, and in that it is chosen from the polypeptides having the following sequences: SEQ. ID No. 488, SEQ ID No. 489, SEQ ID No. 571, SEQ ID No. 572, SEQ No. 573, SEQ ID No. 716 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments that is cysteine-rich and contains RGD sequence, and in that it is chosen from the polypeptides having the following sequence: SEQ ID No. 144 and one of its representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* outer membrane polypeptide that contains cysteines in their first 30 amino acids and also contain an RGD sequence, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 101, SEQ ID No. 122, SEQ ID No. 308, SEQ ID No. 488, SEQ ID No. 489, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 651, SEQ ID No. 679, SEQ ID No. 680, SEQ ID No. 705, SEQ ID No. 716, SEQ ID No. 763, SEQ ID No. 870, SEQ ID No. 878, SEQ ID No. 879, SEQ ID No. 995, SEQ ID No. 1028, SEQ ID No. 1029, SEQ ID No. 1176, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments that contains RGD sequences homologous to *Chlamydia pneumoniae* polypeptides containing RGD sequences, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 28, SEQ ID No. 101, SEQ ID No. 125, SEQ ID No. 155, SEQ ID No. 156, SEQ ID No. 286, SEQ ID No. 571, SEQ ID No. 572, SEQ ID No. 573, SEQ ID No. 763, SEQ ID No. 870, and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a *Chlamydia trachomatis* Type III or non-Type III secreted polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 180, SEQ ID No. 181, SEQ ID No. 207, SEQ ID No. 208, SEQ ID No. 372, SEQ ID No. 391, SEQ ID No. 399, SEQ ID No. 477, SEQ ID No. 486, SEQ ID No. 749, SEQ ID No. 758, SEQ ID No. 819, SEQ ID No. 878, SEQ ID No. 888, SEQ ID No. 896, SEQ ID No. 897, SEQ ID No. 900, SEQ ID No. 902, SEQ ID No. 923, SEQ ID No. 1015, SEQ ID No. 1018, SEQ ID No. 1059, SEQ ID No. 1060, SEQ ID No. 1069, SEQ ID No. 1071, SEQ ID No. 1073, SEQ ID No. 1076, SEQ ID No. 1189, and their representative fragments.

is a *Chlamydia trachomatis* cell wall anchored surface polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences: SEQ ID No. 662, SEQ ID No. 681, SEQ ID No. 1182, SEQ ID No. 1192, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments not found in Chlamydia pneumoniae (Blastp P>e<sup>-10</sup>) and in that it is chosen from the polypeptides having the following sequences: SEQ ID No.2, SEQ ID No. 18, SEQ ID No. 60, SEQ ID No. 66, SEQ ID No. 67, SEQ ID No.68, SEQ ID No. 69, SEQ ID No. 70, SEQ ID No. 81, SEQ ID No. 89, SEQ ID No. 107, SEQ ID 10 No.108, SEQ ID No. 109, SEQ ID No.134, SEQ ID No. 147, SEQ ID No.191, SEQ ID No. 194, SEQ ID No. 216, SEQ ID No. 217, SEQ ID No. 218, SEQ ID No. 219, SEQ ID No. 220, SEQ ID No. 221, SEQ ID No. 222, SEQ ID No. 222, SEQ ID No. 223, SEQ ID No. 224, SEQ ID No. 225, SEQ ID No. 228, SEQ ID No. 235, SEQ ID No. 257, SEQ ID No. 276, SEQ ID No. 277, SEQ ID No. 278, SEQ ID No. 279, SEQ ID No. 280, SEQ ID No. 281, SEQ ID No. 282, SEQ ID No. 283, SEQ ID No. 284, 15 SEQ ID No. 285, SEQ ID No. 289, SEQ ID No.291, SEQ ID No. 298, SEQ ID No. 284, SEQ ID No. 313, SEQ ID No. 314, SEQ ID No. 315, SEQ ID No. 316, SEQ ID No. 334, SEQ ID No. 335, SEQ ID No. 336, SEQ ID No. 337, SEQ ID No. 338, SEQ ID No. 339, SEQ ID No. 340, SEQ ID No. 381, SEQ ID No. 393, SEQ ID No. 413, SEQ ID No. 418, SEQ ID No. 419, SEQ ID No. 419, SEQ ID No. 420, SEQ ID No. 421, SEQ ID No. 422, SEQ ID No. 423, SEQ ID No. 436, SEQ ID No. 460, SEQ ID 20 No. 475, SEQ ID No. 476, SEQ ID No. 480, SEQ ID No. 485, SEQ ID No. 487, SEQ ID No.491, SEQ ID No. 492, SEQ ID No. 493, SEQ ID No. 494, SEQ ID No. 496, SEQ ID No. 500, SEQ ID No. 504, SEQ ID No. 514, SEQ ID No. 527, SEQ ID No. 559, SEQ ID No. 569, SEQ ID No. 570, SEQ ID No. 575, SEQ ID No. 580, SEQ ID No. 582, SEQ ID No. 593, SEQ ID No. 598, SEQ ID No.632, SEQ ID No.640, SEQ ID No.651, SEQ ID No.671, SEQ ID No. 690, SEQ ID No. 694, ID No. 698, SEQ ID 25 No. 710, SEQ ID No. 722, SEQ ID No. 723, SEQ ID No. 724, SEQ ID No. 770, SEQ ID No. 771, SEQ ID No. 782, SEQ ID No. 783, SEQ ID No. 784, SEQ ID No. 790, SEQ ID No. 795, SEQ ID No. 798, SEQ ID No. 805, SEQ ID No. 810, SEQ ID No. 817, SEQ ID No. 829, SEQ ID No. 830, SEQ ID No. 864, SEQ ID No. 866, SEQ ID No. 876, SEQ ID No. 887, SEQ ID No. 892, SEQ ID No. 899, SEQ ID No. 913, SEQ ID No. 921, SEQ ID No. 933, SEQ ID No. 938, SEQ ID No. 949, SEQ ID No. 30 956, SEQ ID No. 1010, SEQ ID No. 1017, SEQ ID No. 1018, SEQ ID No. 1027, SEQ ID No. 1030, SEQ ID No. 1037, SEQ ID No. 1038, SEQ ID No. 1047, SEQ ID No. 1072, SEQ ID No. 1074, SEQ ID No. 1075, SEQ ID No. 1078, SEQ ID No. 1079, SEQ ID No. 1081, SEQ ID No. 1083, SEQ ID No. 1084, SEQ ID No. 1087, SEQ ID No. 1088, SEQ ID No. 1089, SEQ ID No. 1091, SEQ ID No. 1092, SEQ ID No. 1094, SEQ ID No. 1095, SEQ ID No. 1096, SEQ ID No. 1098, SEQ ID No. 1104, SEQ 35 ID No. 1105, SEQ ID No. 1106, SEQ ID No. 1108, SEQ ID No. 1110, SEQ ID No. 1114, SEQ ID No.

1115, SEQ ID No. 1116, SEQ ID No. 1117, SEQ ID No. 1119, SEQ ID No. 1128, SEQ ID No. 1132, SEQ ID No. 1133, SEQ ID No. 1135, SEQ ID No. 1136, SEQ ID No. 1139, SEQ ID No. 1140, SEQ ID No. 1141, SEQ ID No. 1142, SEQ ID No. 1144, SEQ ID No. 1148, SEQ ID No. 1151, SEQ ID No. 1155, SEQ ID No. 1157, SEQ ID No. 1159, SEQ ID No. 1161, SEQ ID No. 1162, SEQ ID No. 1165, SEQ ID No. 1166, SEQ ID No. 1167, SEQ ID No. 1168, SEQ ID No. 1169, SEQ ID No. 1171, SEQ ID No. 1172, SEQ ID No. 1173, SEQ ID No. 1174, SEQ ID No. 1175, SEQ ID No. 1176, SEQ ID No. 1177, SEQ ID No. 1178, SEQ ID No. 1180, SEQ ID No. 1181, SEQ ID No. 1183, SEQ ID No. 1184, SEQ ID No. 1186, SEQ ID No. 1187, SEQ ID No. 1188, SEQ ID No. 1192, SEQ ID No. 1194, SEQ ID No. 1197, and their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 10; SEQ ID No. 44; SEQ ID No. 45; SEQ ID No. 46; SEQ ID No. 47; SEQ ID No. 93; 15 SEQ ID No. 101; SEQ ID No. 102; SEQ ID No. 103; SEQ ID No. 106; SEQ ID No. 107; SEQ ID No. 120; SEQ ID No. 121; SEQ ID No. 130; SEQ ID No. 135; SEQ ID No. 140; SEQ ID No. 143; SEQ ID No. 144; SEQ ID No. 145; SEQ ID No. 158; SEQ ID No. 159; SEQ ID No. 160; SEQ ID No. 161; SEQ ID No. 192; SEQ ID No. 193; SEQ ID No. 196; SEQ ID No. 197; SEQ ID No. 198; SEQ ID No. 199; SEQ ID No. 227; SEQ ID No. 229; 20 SEQ ID No. 236; SEQ ID No. 236; SEQ ID No. 239; SEQ ID No. 243; SEQ ID No. 245; SEQ ID No. 264; SEQ ID No. 265; SEQ ID No. 297; SEQ ID No. 331; SEQ ID No. 333; SEQ ID No. 359; SEQ ID No. 360; SEQ ID No. 374; SEQ ID No. 404; SEQ ID No. 405; SEQ ID No. 405; SEQ ID No. 410; SEQ ID No. 415; SEQ ID No. 415; SEQ ID No. 416; SEQ ID No. 417; SEQ ID No. 432; SEQ ID No. 460; SEQ ID No. 461; SEQ ID No. 462; 25 SEQ ID No. 495; SEQ ID No. 513; SEQ ID No. 515; SEQ ID No. 566; SEQ ID No. 566; SEQ ID No. 566; SEQ ID No. 589; SEQ ID No. 613; SEQ ID No. 645; SEQ ID No. 646; SEQ ID No. 647; SEQ ID No. 652; SEQ ID No. 653; SEQ ID No. 654; SEQ ID No. 672; SEQ ID No. 673; SEQ ID No. 674; SEQ ID No. 682; SEQ ID No. 684; SEQ ID No. 692; SEQ ID No. 700; SEQ ID No. 725; SEQ ID No. 801; SEQ ID No. 802; SEQ ID No. 835; 30 SEQ ID No. 836; SEQ ID No. 837; SEQ ID No. 860; SEQ ID No. 861; SEQ ID No. 862; SEQ ID No. 863; SEQ ID No. 869; SEQ ID No. 869; SEQ ID No. 925; SEQ ID No. 964; SEQ ID No. 983 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the intermediate metabolism of nucleotides or nucleic acids, and in that it is

chosen from the polypeptides having the following sequences:

```
SEQ ID No. 142;
                      SEQ ID No. 142;
                                         SEQ ID No. 169;
                                                            SEQ ID No. 256;
                                                                               SEQ ID No. 268;
   SEQ ID No. 325;
                      SEQ ID No. 352;
                                         SEQ ID No. 366;
                                                            SEQ ID No. 435;
                                                                               SEQ ID No. 444;
   SEQ ID No. 528;
                      SEQ ID No. 529;
                                         SEQ ID No. 530;
                                                            SEQ ID No. 548;
                                                                               SEQ ID No. 549;
5 SEQ ID No. 601;
                      SEQ ID No. 602;
                                         SEQ ID No. 617;
                                                            SEQ ID No. 619;
                                                                               SEQ ID No. 644;
   SEQ ID No. 745; SEQ ID No. 971; SEQ ID No. 972; SEQ ID No. 1023 and one of their representative
   fragments.
```

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of nucleic acids, and in that it is chosen from the polypeptides having the following sequences:

```
SEQ ID No. 5; SEQ ID No. 12; SEQ ID No. 82; SEQ ID No. 96; SEQ ID No. 97; SEQ ID No. 98;
     SEQ ID No. 99;
                       SEQ ID No. 100;
                                          SEQ ID No. 105;
                                                            SEQ ID No. 118;
                                                                               SEQ ID No. 136;
     SEQ ID No. 137;
                       SEQ ID No. 163;
                                          SEQ ID No. 190;
                                                             SEQ ID No. 204;
                                                                               SEQ ID No. 259;
    SEQ ID No. 260;
                       SEQ ID No. 262;
                                          SEQ ID No. 290;
                                                             SEQ ID No. 300;
                                                                               SEQ ID No. 301;
     SEQ ID No. 302;
                       SEQ ID No. 387;
                                          SEQ ID No. 427;
                                                             SEQ ID No. 434;
                                                                               SEQ ID No. 441;
    SEQ ID No. 444;
                       SEQ ID No. 471;
                                          SEQ ID No. 595;
                                                             SEQ ID No. 596;
                                                                               SEQ ID No. 597;
    SEQ ID No. 599;
                       SEQ ID No. 600;
                                          SEQ ID No. 605;
                                                            SEQ ID No. 612;
                                                                               SEQ ID No. 624;
    SEQ ID No. 625;
                       SEQ ID No. 650;
                                          SEQ ID No. 657;
                                                            SEQ ID No. 658;
                                                                               SEQ ID No. 702;
    SEQ ID No. 703;
                       SEQ ID No. 704;
                                          SEQ ID No. 708;
                                                            SEQ ID No. 719;
                                                                               SEQ ID No. 766;
    SEQ ID No. 767;
                       SEQ ID No. 775;
                                          SEQ ID No. 779;
                                                            SEQ ID No. 787;
                                                                               SEQ ID No. 788;
    SEQ ID No. 794;
                       SEQ ID No. 841;
                                          SEQ ID No. 842;
                                                            SEQ ID No. 883;
                                                                               SEQ ID No. 884;
    SEQ ID No. 907;
                       SEQ ID No. 918;
                                          SEQ ID No. 924;
                                                            SEQ ID No. 928;
                                                                               SEQ ID No. 929;
    SEQ ID No. 962;
                       SEQ ID No. 962;
                                          SEQ ID No. 963;
                                                            SEQ ID No. 969;
                                                                               SEQ ID No. 970;
25 SEQ ID No. 975; SEQ ID No. 979; SEQ ID No. 995; SEQ ID No. 1031; SEQ ID No. 1032 and one of
    their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of amino acids or polypeptides, and in that it is chosen from the polypeptides having the following sequences:

```
SEQ ID No. 27; SEQ ID No. 41; SEQ ID No. 55; SEQ ID No. 56; SEQ ID No. 57; SEQ ID No. 59;
    SEQ ID No. 62; SEQ ID No. 63; SEQ ID No. 64; SEQ ID No. 65; SEQ ID No. 119; SEQ ID No. 132;
    SEQ ID No. 240;
                       SEQ ID No. 241;
                                          SEQ ID No. 277;
                                                            SEQ ID No. 278;
                                                                               SEQ ID No. 279;
    SEQ ID No. 382;
                       SEQ ID No. 406;
                                          SEQ ID No. 428;
                                                            SEQ ID No. 442;
                                                                               SEQ ID No. 446;
35 SEQ ID No. 447;
                       SEQ ID No. 453;
                                          SEQ ID No. 454;
                                                            SEQ ID No. 541;
                                                                               SEQ ID No. 542;
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SEQ ID No. 591; SEQ ID No. 608; SEQ ID No. 609; SEQ ID No. 610; SEQ ID No. 618; SEQ ID No. 648; SEQ ID No. 649; SEQ ID No. 660; SEQ ID No. 661; SEQ ID No. 677; SEQ ID No. 717; SEQ ID No. 765; SEQ ID No. 797; SEQ ID No. 871; SEQ 1D No. 875; SEQ ID No. 920; SEQ ID No. 922; SEQ ID No. 937; SEQ ID No. 998; SEQ ID No. 1020; SEQ ID No. 1021; SEQ ID No. 1034; SEQ ID No. 1044; SEQ ID No. 1046; SEQ ID No. 1049 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of polypeptides, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 21; SEQ ID No. 22; SEQ ID No. 23; SEQ ID No. 24; SEQ ID No. 25; SEQ ID No. 26; SEQ ID No. 75; SEQ ID No. 84; SEQ ID No. 86; SEQ ID No. 92; SEQ ID No. 133; SEQ ID No. 151; SEQ ID No. 152; SEQ ID No. 157; SEQ ID No. 179; SEQ ID No. 209; SEQ ID No. 307; SEQ ID No. 326; SEQ ID No. 343; SEQ ID No. 344; SEQ ID No. 345; SEQ ID No. 371; 15 SEQ ID No. 429; SEQ ID No. 519; SEQ ID No. 557; SEQ ID No. 586; SEQ ID No. 587; SEQ ID No. 630; SEQ ID No. 656; SEQ ID No. 706; SEQ ID No. 707; SEQ ID No. 730; SEQ ID No. 751; SEQ ID No. 752; SEQ ID No. 786; SEQ ID No. 847; SEQ ID No. 885; SEQ ID No. 923; SEQ ID No. 978; SEQ ID No. 1039; SEQ ID No. 1048 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the metabolism of fatty acids, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 4; SEQ ID No. 15; SEQ ID No. 16; SEQ ID No. 141; SEQ ID No. 173; SEQ ID No. 205; 25 SEQ ID No. 205; SEQ ID No. 206; SEQ ID No. 207; SEQ ID No. 208; SEQ ID No. 312; SEQ ID No. 355; SEQ ID No. 415; SEQ ID No. 550; SEQ ID No. 558; SEQ ID No. 560; SEQ ID No. 561; SEQ ID No. 574; SEQ ID No. 574; SEQ ID No. 577; SEQ ID No. 578; SEQ ID No. 590; SEQ ID No. 614; SEQ ID No. 772; SEQ ID No. 808; SEQ ID No. 809; SEQ ID No. 904; SEQ ID No. 905; SEQ ID No. 905; SEQ ID No. 933; SEQ ID No. 934;

30 SEQ ID No. 934; SEQ ID No. 936 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the synthesis of the wall, and in that it is chosen from the polypeptides having the following sequences:

35 SEQ ID No. 87; SEQ ID No. 196; SEQ ID No. 242; SEQ ID No. 269; SEQ ID No. 628;

SEQ ID No. 629; SEQ ID No. 634; SEQ ID No. 635; SEQ ID No. 637; SEQ ID No. 638; SEQ ID No. 1019 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* polypeptide or one of its representative fragments which is involved in the transcription, translation and/or maturation process, and in that it is chosen from the polypeptides having the following sequences:

```
SEQ ID No. 112;
                   SEQ ID No. 113;
                                      SEQ ID No. 332;
                                                         SEQ ID No. 212;
                                                                           SEQ ID No. 213;
SEQ ID No. 350;
                   SEQ ID No. 362;
                                      SEQ ID No. 363;
                                                         SEQ ID No. 364;
                                                                           SEQ ID No. 407:
SEQ ID No. 451;
                   SEQ ID No. 546;
                                      SEQ ID No. 643;
                                                         SEQ ID No. 744;
                                                                           SEQ ID No. 746;
SEQ ID No. 833;
                   SEQ ID No. 868;
                                     SEQ ID No. 981;
                                                       SEQ ID No. 982;
                                                                          SEQ ID No. 1003;
SEQ ID No. 1011; SEQ ID No. 1042 and one of their representative fragments.
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Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* ribosomal polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

```
15 SEQ ID No. 114;
                       SEQ ID No. 115;
                                         SEQ ID No. 116;
                                                           SEQ ID No. 328;
                                                                              SEQ ID No. 361;
    SEQ ID No. 375;
                       SEQ ID No. 445;
                                         SEQ ID No. 543;
                                                           SEQ ID No. 584;
                                                                              SEQ ID No. 585;
    SEQ ID No. 743;
                       SEQ ID No. 813;
                                         SEQ ID No. 941;
                                                           SEQ ID No. 942;
                                                                             SEQ ID No. 944;
    SEQ ID No. 946:
                       SEQ ID No. 947;
                                         SEQ ID No. 948;
                                                           SEQ ID No. 950;
                                                                             SEQ ID No. 951;
    SEQ ID No. 952;
                                         SEQ ID No. 954;
                       SEQ ID No. 953;
                                                           SEQ ID No. 955;
                                                                             SEQ ID No. 955;
20 SEQ ID No. 957;
                      SEQ ID No. 958;
                                        SEQ ID No. 960;
                                                          SEQ ID No. 961;
                                                                            SEQ ID No. 1040;
    SEQ ID No. 1041; SEQ ID No. 1043; SEQ ID No. 1063; SEQ ID No. 1064 and one of their
    fragments.
```

Preferably, the invention also relates to a polypeptide according to the invention, characterized in that it is a *Chlamydia trachomatis* transport polypeptide or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

```
SEQ ID No. 6; SEQ ID No. 50; SEQ ID No. 51; SEQ ID No. 80; SEQ ID No. 125; SEQ ID No. 126;
    SEQ ID No. 128;
                       SEQ ID No. 129;
                                          SEQ ID No. 215;
                                                            SEQ ID No. 246;
                                                                               SEQ ID No. 248;
    SEQ ID No. 249;
                       SEQ ID No. 251;
                                          SEQ ID No. 252;
                                                            SEQ ID No. 253;
                                                                               SEQ ID No. 255;
    SEQ ID No. 271;
                       SEQ ID No. 275;
                                          SEQ ID No. 293;
                                                            SEQ ID No. 309;
                                                                               SEQ ID No. 323;
30 SEQ ID No. 324;
                       SEQ ID No. 398;
                                          SEQ ID No. 401:
                                                            SEQ ID No. 449;
                                                                               SEQ ID No. 511;
    SEQ ID No. 512;
                       SEQ ID No. 564:
                                          SEQ ID No. 565;
                                                            SEQ ID No. 667;
                                                                               SEQ ID No. 679;
    SEQ ID No. 680:
                       SEQ ID No. 711;
                                          SEQ ID No. 712;
                                                            SEQ ID No. 713;
                                                                               SEQ ID No. 714;
    SEQ ID No. 715;
                       SEQ ID No. 730;
                                          SEQ ID No. 731;
                                                            SEQ ID No. 736;
                                                                               SEQ ID No. 737;
    SEQ ID No. 738;
                       SEQ ID No. 870;
                                          SEQ ID No. 908;
                                                            SEQ ID No. 919;
                                                                               SEQ ID No. 977;
35 SEQ ID No. 987;
                       SEQ ID No. 988;
                                          SEQ ID No. 992;
                                                            SEQ ID No. 993;
                                                                               SEQ ID No. 994;
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35 SEQ ID No. 433;

SEQ ID No. 1028; SEQ ID No. 1029 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments which is involved in the virulence process, and in that it is chosen from the polypeptides having the 5 following sequences:

SEQ ID No. 20; SEQ ID No. 815; SEQ ID No. 816; SEQ ID No. 898; SEQ ID No. 1059; SEQ ID No. 1060 and one of their representative fragments.

Preferably, the invention relates to a polypeptide according to the invention, characterized in that it is a Chlamydia trachomatis polypeptide or one of its representative fragments 10 which is involved in the secretory system and/or which is secreted, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 758; SEQ ID No. 888; SEQ ID No. 889; SEQ ID No. 890; SEQ ID No. 891; SEQ ID No. 896; SEQ ID No. 897; SEQ ID No. 898 and one of their representative fragments.

The secreted polypeptides, including the Type III and other, non-Type III secreted polypeptides, of the present invention, as well as the corresponding nucleotide sequences, may be detected by techniques known to persons skilled in the art, such as for example the techniques using cloning combined with vectors allowing the expression of the said polypeptides fused to export markers such as the luc gene for luciferase or the PhoA gene for alkaline phosphatase.

Preferably, the invention relates to a polypeptide according to the invention, 20 characterized in that it is a polypeptide specific to Chlamydiae or one of its representative fragments, and in that it is chosen from the polypeptides having the following sequences:

SEQ ID No. 22; SEQ ID No. 29; SEQ ID No. 31; SEQ ID No. 32; SEQ ID No. 34; SEQ ID No. 35; SEQ ID No. 39; SEQ ID No. 40; SEQ ID No. 43; SEQ ID No. 48; SEQ ID No. 49; SEQ ID No. 50; SEQ ID No. 52; SEQ ID No. 53; SEQ ID No. 54; SEQ ID No. 72; SEQ ID No. 77; SEQ ID No. 78; 25 SEQ ID No. 87; SEQ ID No. 90; SEQ ID No. 95; SEQ ID No. 108; SEQ ID No. 110; SEQ ID No. 111; SEQ ID No. 122; SEQ ID No. 123; SEQ ID No. 124; SEQ ID No. 127; SEQ ID No. 138; SEQ ID No. 144; SEQ ID No. 146; SEQ ID No. 153; SEQ ID No. 155; SEQ ID No. 164; SEQ ID No. 166; SEQ ID No. 175; SEQ ID No. 182; SEQ ID No. 184; SEQ ID No. 186; SEQ ID No. 187; SEQ ID No. 188; SEQ ID No. 202; SEQ ID No. 210; 30 SEQ ID No. 247; SEQ ID No. 258; SEQ ID No. 266; SEQ ID No. 267; SEQ ID No. 270; SEQ ID No. 273; SEQ ID No. 274; SEQ ID No. 295; SEQ ID No. 296; SEQ ID No. 305; SEQ ID No. 306; SEQ ID No. 318; SEQ ID No. 309; SEQ ID No. 319; SEQ ID No. 322; SEQ ID No. 326; SEQ ID No. 342; SEQ ID No. 357; SEQ ID No. 376; SEQ ID No. 379; SEQ ID No. 380; SEQ ID No. 388;

SEQ ID No. 390;

SEQ ID No. 443;

SEQ ID No. 438;

SEQ ID No. 400;

SEQ ID No. 456;

SEQ ID No. 431;

SEQ ID No. 457;

	SEQ ID No. 458;	SEQ ID No. 464;	SEQ ID No. 468;	SEQ ID No. 470;	SEQ ID No. 473;
	SEQ ID No. 486;	SEQ ID No. 489;	SEQ ID No. 497;	SEQ ID No. 501;	SEQ ID No. 503;
	SEQ ID No. 504;	SEQ ID No. 508;	SEQ ID No. 512;	SEQ ID No. 521;	SEQTD No. 522;
	SEQ ID No. 523;	SEQ ID No. 524;	SEQ ID No. 533;	SEQ ID No. 535;	SEQ ID No. 536;
5	SEQ ID No. 537;	SEQ ID No. 538;	SEQ ID No. 539;	SEQ ID No. 540;	SEQ ID No. 554;
	SEQ ID No. 563;	SEQ ID No. 572;	SEQ ID No. 579;	SEQ ID No. 595;	SEQ ID No. 603;
	SEQ ID No. 604;	SEQ ID No. 606;	SEQ ID No. 607;	SEQ ID No. 615;	SEQ ID No. 616;
	SEQ ID No. 622;	SEQ ID No. 641;	SEQ ID No. 642;	SEQ ID No. 659;	SEQ ID No. 668;
	SEQ ID No. 670;	SEQ ID No. 693;	SEQ ID No. 695;	SEQ ID No. 696;	SEQ ID No. 699;
10	SEQ ID No. 703;	SEQ ID No. 704;	SEQ ID No. 716;	SEQ ID No. 726;	SEQ ID No. 728;
	SEQ ID No. 739;	SEQ ID No. 742;	SEQ ID No. 747;	SEQ ID No. 750;	SEQ ID No. 751;
	SEQ ID No. 755;	SEQ ID No. 757;	SEQ ID No. 759;	SEQ ID No. 761;	SEQ ID No. 762;
	SEQ ID No. 763;	SEQ ID No. 764;	SEQ ID No. 773;	SEQ ID No. 780;	SEQ ID No. 781;
	SEQ ID No. 789;	SEQ ID No. 800;	SEQ ID No. 803;	SEQ ID No. 804;	SEQ ID No. 818;
15	SEQ ID No. 820;	SEQ ID No. 822;	SEQ ID No. 823;	SEQ ID No. 824;	SEQ ID No. 827;
	SEQ ID No. 828;	SEQ ID No. 839;	SEQ ID No. 849;	SEQ ID No. 850;	SEQ ID No. 851;
	SEQ ID No. 852;	SEQ ID No. 855;	SEQ ID No. 856;	SEQ ID No. 857;	SEQ ID No. 858;
	SEQ ID No. 859;	SEQ ID No. 860;	SEQ ID No. 861;	SEQ ID No. 862;	SEQ ID No. 863;
	SEQ ID No. 865;	SEQ ID No. 868;	SEQ ID No. 869;	SEQ ID No. 870;	SEQ ID No. 871;
20	SEQ ID No. 872;	SEQ ID No. 873;	SEQ ID No. 874;	SEQ ID No. 875;	SEQ ID No. 877;
	SEQ ID No. 878;	SEQ ID No. 880;	SEQ ID No. 882;	SEQ ID No. 884;	SEQ ID No. 886;
	SEQ ID No. 893;	SEQ ID No. 901;	SEQ ID No. 906;	SEQ ID No. 910;	SEQ ID No. 912;
	SEQ ID No. 915;	SEQ ID No. 916;	SEQ ID No. 917;	SEQ ID No. 926;	SEQ ID No. 929;
	SEQ ID No. 933;	SEQ ID No. 965;	SEQ ID No. 967;	SEQ ID No. 968;	SEQ ID No. 984;
25	SEQ ID No. 986;	SEQ ID No. 989;	SEQ ID No. 990;	SEQ ID No. 996;	SEQ ID No. 997;
	SEQ ID No. 1001;	SEQ ID No. 1002;	SEQ ID No. 1013;	SEQ ID No. 1016;	SEQ ID No. 1031;
	SEQ ID No. 1033;	SEQ ID No. 1035;	SEQ ID No. 1049;	SEQ ID No. 1051;	SEQ ID No. 1052;
	SEQ ID No. 1054;	SEQ ID No. 1056;	SEQ ID No. 1057;	SEQ ID No. 1058;	SEQ ID No. 1062;
	SEQ ID No. 1070;	SEQ ID No. 1071; SE	EQ ID No. 1073 and o	one of their represents	ative fragments.
30	ln e	reneral in the present	invantion the forces	innat annu de la tal	

ln general, in the present invention, the functional group to which a polypeptide of the invention belongs, as well as its corresponding nucleotide sequence, may be determined either by comparative analogy with sequences already known, or by the use of standard techniques of biochemistry, of cytology combined with the techniques of genetic engineering such as immunoaffinity, localization by immunolabelling, differential extraction, measurement of enzymatic activity, study of the activity inducing or repressing expression or the study of expression in *E. coli*.

It is clearly understood, on the one hand, that, in the present invention, the nucleotide sequences (ORF) and the amino acid sequences (SEQ ID No. 2 to SEQ ID No. 1197) which are listed by functional group, are not exhaustive within the group considered. Moreover, it is also clearly understood that, in the present invention, a nucleotide sequence (ORF) or an amino acid sequence mentioned within a given functional group may also be part of another group taking into account, for example, the interrelationship between the groups listed. Accordingly, and as an example of this interrelationship, an exported and/or secreted polypeptide as well as its coding nucleotide sequence may also be involved in the *Chlamydia trachomatis* virulence process by modifying the defense mechanism of the infected host cell, or a transmembrane polypeptide or its coding nucleotide sequence is also part of the polypeptides or coding nucleotide sequences of the cellular envelope.

The subject of the present invention is also the nucleotide and/or polypeptide sequences according to the invention, characterized in that the said sequences are recorded on a medium, called recording medium, whose type and nature facilitate the reading, the analysis and the exploitation of the said sequences. These media may of course also contain other information extracted from the present invention, such as in particular the analogies with already known sequences, such as those mentioned in Table 1 of the present description, and/or may contain, in addition, information relating to the nucleotide and/or polypeptide sequences of other microorganisms so as to facilitate the comparative analysis and the exploitation of the results obtained.

Among these recording media, computer-readable media, such as magnetic, optical, electrical and hybrid media such as, for example, floppy disks, CD-ROMs or recording cassettes, are preferred in particular.

The invention also relates to nucleotide sequences which can be used as primer or probe, characterized in that the said sequences are chosen from the nucleotide sequences according to the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, as primer or probe, for the detection and/or amplification of nucleic acid sequences.

The nucleotide sequences according to the invention may thus be used to amplify nucleotide sequences, in particular by the PCR technique (polymerase chain reaction) (Erlich, 1989; Innis et al., 1990; Rolfs et al., 1991, and White et al., 1997).

These oligodeoxyribonucleotide or oligoribonucleotide primers correspond to representative nucleotide fragments, and are advantageously at least 8 nucleotides, preferably at least 12 nucleotides, 15 nucleotides and still more preferably at least 20 nucleotides long.

Other techniques for amplifying the target nucleic acid may be advantageously used as alternatives to PCR.

The nucleotide sequences of the invention, in particular the primers according to the

invention, may also be used in other methods for amplifying a target nucleic acid, such as:

- the TAS (Transcription-based Amplification System) technique described by Kwoh et al. in 1989;
- the 3SR (Self-Sustained Sequence Replication) technique described by Guatelli et al. in 1990;
- 5 the NASBA (Nucleic Acid Sequence Based Amplification) technique described by Kievitis et al. in 1991;
  - the SDA (Strand Displacement Amplification) technique (Walker et al., 1992);
  - the TMA (Transcription Mediated Amplification) technique.

The polynucleotides of the invention may also be used in techniques for amplifying or for modifying the nucleic acid serving as probe, such as:

- the LCR (Ligase Chain Reaction) technique described by Landegren et al. in 1988 and perfected by Barany et al. in 1991, which uses a thermostable ligase;
- the RCR (Repair Chain Reaction) technique described by Segev in 1992;
- the CPR (Cycling Probe Reaction) technique described by Duck et al. in 1990;
- the Q-beta-replicase amplification technique described by Miele et al. in 1983 and perfected in particular by Chu et al. in 1986, Lizardi et al. in 1988, and then by Burg et al. as well as by Stone et al. in 1996.

The invention also relates to the nucleotide sequences of fragments which can be obtained by amplification with the aid of at least one primer according to the invention. The present invention encompasses both hybridization probes and primers. In general, the complementary probes should be of the length sufficient to form a stable hybrid complex with the target sequences. Primers, while complementary to the target sequences need not form stable hybridization complexes with the target sequences alone. Rather, primers form stable complexes with the target sequences in the presence of polymerase to permit extension of the primer.

In the case where the target polynucleotide to be detected is possibly an RNA, for example an mRNA, it will be possible to use, prior to the use of an amplification reaction with the aid of at least one primer according to the invention or to the use of a method of detection with the aid of at least one probe of the invention, a reverse transcriptase-type enzyme so as to obtain a cDNA from the RNA contained in the biological sample. The cDNA obtained will then serve as target for the primer(s) or the probe(s) used in the amplification or detection method according to the invention.

The detection probe will be chosen so that it hybridizes with the target sequence or the amplicon generated from the target sequence. Such a detection probe will advantageously have as sequence a sequence of at least 12 nucleotides, 15 nucleotides, in particular of at least 20 nucleotides, and preferably at least 100 nucleotides.

The invention also comprises the nucleotide sequences which can be used as probe or

primer according to the invention, characterized in that they are labelled with a radioactive compound or with a nonradioactive compound.

The nonlabelled nucleotide sequences may be used directly as probes or primers; however, the sequences are generally labelled with a radioactive element (<sup>32</sup>P, <sup>35</sup>S, <sup>3</sup>H, <sup>125</sup>I) or with a nonradioactive molecule (biotin, acetylaminofluorene, digoxigenin, 5-bromo-deoxyuridine, fluorescein) so as to obtain probes which can be used in numerous applications.

Examples of nonradioactive labelling of nucleotide sequences are described, for example, in French patent No. 78,10975 or by Urdea et al. or by Sanchez-Pescador et al. in 1988.

In the latter case, one of the labelling methods described in patents FR-2 422 956 and FR-2 518 755 may also be used.

The invention also relates to the nucleotide sequences of fragments which can be obtained by hybridization with the aid of at least one probe according to the invention.

The hybridization technique may be performed in various ways (Matthews et al., 1988). The most common method consists in immobilizing the nucleic acid extracted from C. trachomatis cells on a support (such as nitrocellulose, nylon, polystyrene) and in incubating, under well-defined conditions, the target nucleic acid immobilized with the probe. After hybridization, the excess probe is removed and the hybrid molecules formed are detected by the appropriate method (measurement of the radioactivity, of the fluorescence or of the enzymatic activity linked to the probe).

The invention also comprises the nucleotide sequences according to the invention, characterized in that they are covalently or noncovalently immobilized on a support.

According to another advantageous embodiment of the nucleic sequences according to the invention, the latter may be used immobilized on a support and may thus serve to capture, through specific hybridization, the target nucleic acid obtained from the biological sample to be tested. If necessary, the solid support is separated from the sample and the hybridization complex formed between the so-called capture probe and the target nucleic acid is then detected by means of a second probe, called detection probe, labelled with an easily detectable element.

The nucleotide sequences according to the invention may also be used in new analytical systems, DNA chips, which allow sequencing, the study of mutations and of the expression of genes, and which are currently of interest given their very small size and their high capacity in terms of number of analyses.

The principle of the operation of these chips is based on molecular probes, most often oligonucleotides, which are attached onto a miniaturized surface, generally of the order of a few square centimetres. During an analysis, a sample containing fragments of a target nucleic acid to be analysed, for example DNA or RNA labelled, for example, after amplification, is deposited onto the

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DNA chip in which the support has been coated beforehand with probes. Bringing the labelled target sequences into contact with the probes leads to the formation, through hybridization, of a duplex according to the rule of pairing defined by J.D. Watson and F. Crick. After a washing step, analysis of the surface of the chip allows the effective hybridizations to be located by means of the signals emitted by the labels tagging the target. A hybridization fingerprint results from this analysis which, by appropriate computer processing, will make it possible to determine information such as the presence of specific fragments in the sample, the determination of sequences and the presence of mutations.

The chip consists of a multitude of molecular probes, precisely organized or arrayed on a solid support whose surface is miniaturized. It is at the centre of a system where other elements (imaging system, microcomputer) allow the acquisition and interpretation of a hybridization fingerprint.

The hybridization supports are provided in the form of flat or porous surfaces (pierced with wells) composed of various materials. The choice of a support is determined by its physicochemical properties, or more precisely, by the relationship between the latter and the conditions under which the support will be placed during the synthesis or the attachment of the probes or during the use of the chip. It is therefore necessary, before considering the use of a particular support (R.S. Matson et al., 1994), to consider characteristics such as its stability to pH, its physical strength, its reactivity and its chemical stability as well as its capacity to nonspecifically bind nucleic acids. Materials such as glass, silicon and polymers are commonly used. Their surface is, in a first step, called "functionalization", made reactive towards the groups which it is desired to attach thereon. After the functionalization, so-called spacer molecules are grafted onto the activated surface. Used as intermediates between the surface and the probe, these molecules of variable size render unimportant the surface properties of the supports, which often prove to be problematic for the synthesis or the attachment of the probes and for the hybridization.

Among the hybridization supports, there may be mentioned glass which is used, for example, in the method of in situ synthesis of oligonucleotides by photochemical addressing developed by the company Affymetrix (E.L. Sheldon, 1993), the glass surface being activated by silane. Genosensor Consortium (P. Mérel, 1994) also uses glass slides carrying wells 3 mm apart, this support being activated with epoxysilane.

Polymers or silicon may also be mentioned among these hybridization supports. For example, the Andrein Mirzabekov team has developed a chip consisting of polyacrylamide squares polymerized on a silanized glass surface (G. Yershov et al., 1996). Several teams use silicon, in particular the IFOS laboratory of Ecole Centrale of Lyon which uses a silicon semiconductor substrate which is p-doped by introducing it into its crystalline structure atoms whose valency is different from

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that of silicon. Various types of metals, in particular gold and platinum, may also be used as support (Genosensor Consortium (K. Beattie et al., 1993)).

The probes according to the invention may be synthesized directly in situ on the supports of the DNA chips. This in situ synthesis may be carried out by photochemical addressing 5 (developed by the company Affymax (Amsterdam, Holland) and exploited industrially by its subsidiary Affymetrix (United States)) or based on the VLSIPS (very large scale immobilized polymer synthesis) technology (S.P.A. Fodor et al., 1991) which is based on a method of photochemically directed combinatory synthesis and the principle of which combines solid-phase chemistry, the use of photolabile protecting groups and photolithography.

The probes according to the invention may be attached to the DNA chips in various ways such as electrochemical addressing, automated addressing or the use of probe printers (T. Livache et al., 1994; G. Yershov et al., 1996; J. Derisi et al., 1996, and S. Borman, 1996).

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The revealing of the hybridization between the probes of the invention, deposited or synthesized in situ on the supports of the DNA chips, and the sample to be analysed, may be 15 determined, for example, by measurement of fluorescent signals, by radioactive counting or by electronic detection.

The use of fluorescent molecules such as fluorescein constitutes the most common method of labelling the samples. It allows direct or indirect revealing of the hybridization and allows the use of various fluorochromes.

Affymetrix currently provides an apparatus or a scanner designed to read its Gene Chip? chips. It makes it possible to detect the hybridizations by scanning the surface of the chip in confocal microscopy (R.J. Lipshutz et al., 1995). Other methods of detecting fluorescent signals have been tested: coupling of an epifluorescence microscope and a CCD camera (G. Yershov et al., 1996), the use of an optical fibre collecting system (E.L. Sheldon, 1993). A conventional method consists in 25 carrying out an end labelling, with phosphorus 32, of the target sequences, by means of an appropriate apparatus, the Phosphorimager (marketed by Molecular Dynamics). The electronic detection is based on the principle that the hybridization of two nucleic acid molecules is accompanied by physical phenomena which can be quantified under certain conditions (system developed by Ecole Centrale of Lyon and called GEN-FET (GEN field effect transistor)). Genosensor Consortium and the company 30 Beckman Instruments who are developing an electronic chip or Permittivity Chips? may also be mentioned (K. Beattie et al., 1993).

The nucleotide sequences according to the invention may thus be used in DNA chips to carry out the analysis of mutations. This analysis is based on the production of chips capable of analysing each base of a nucleotide sequence according to the invention. It is possible, in particular to this end, to use the microsequencing techniques on a DNA chip. The mutations are detected by

extending immobilized primers which hybridize to the template of sequences analysed, just at the position adjacent to that of the mutated nucleotide to be detected. A single-stranded template, RNA or DNA, of the sequences to be analysed will be advantageously prepared according to conventional methods, from products amplified according to PCR-type techniques. The templates of single-stranded DNA, or of RNA thus obtained are then deposited on the DNA chip, under conditions allowing their specific hybridization to the immobilized primers. A thermostable polymerase, for example Tth or T7 DNA polymerase, specifically extends the 3' end of the immobilized primer with a labelled nucleotide analogue complementary to the nucleotide at the position of the variable site. For example a thermal cycling is performed in the presence of fluorescent dideoxyribonucleotides.

Or The experimental conditions will be adapted in particular to the chips used, to the immobilized primers, to the polymerases used and to the labelling system chosen. One advantage of microsequencing, compared with techniques based on the hybridization of probes, is that it makes it possible to identify all the variable nucleotides with optimal discrimination under homogeneous reaction conditions; used on DNA chips, it allows optimal resolution and specificity for the routine and industrial detection of mutations in multiplex.

The nucleotide sequences according to the invention may also be used in DNA chips to carry out the analysis of the expression of the Chlamydia trachomatis genes. This analysis of the expression of Chlamydia trachomatis genes is based on the use of chips where probes of the invention, chosen for their specificity to characterize a given gene, are present (D.J. Lockhart et al., 1996; D.D. Shoemaker et al., 1996). For the methods of analysis of gene expression using the DNA chips, reference may, for example, be made to the methods described by D.J. Lockhart et al. (1996) and Sosnowsky et al. (1997) for the synthesis of probes in situ or for the addressing and the attachment of previously synthesized probes. The target sequences to be analysed are labelled and in general fragmented into sequences of about 50 to 100 nucleotides before being hybridized onto the chip. After washing as described, for example, by D.J. Lockhart et al. (1996) and application of different electric fields (Sosnowsky et al., 1997), the labelled compounds are detected and quantified, the hybridizations being carried out at least in duplicate. Comparative analyses of the signal intensities obtained with respect to the same probe for different samples and/or for different probes with the same sample, determine the differential expression of RNA or of DNA derived from the sample.

The nucleotide sequences according to the invention may, in addition, be used in DNA chips where other nucleotide probes specific for other microorganisms are also present, and may allow the carrying out of a serial test allowing rapid identification of the presence of a microorganism in a sample.

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to the invention, characterized in that they are immobilized on a support of a DNA chip.

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The DNA chips, characterized in that they contain at least one nucleotide sequence according to the invention, immobilized on the support of the said chip, also form part of the invention.

The said chips will preferably contain several probes or nucleotide sequences of the invention of different length and/or corresponding to different genes so as to identify, with greater certainty, the specificity of the target sequences or the desired mutation in the sample to be analysed.

Accordingly, the analyses carried out by means of primers and/or probes according to the invention, immobilized on supports such as DNA chips, will make it possible, for example, to identify, in samples, mutations linked to variations such as intraspecies variations. These variations may be correlated or associated with pathologies specific to the variant identified and will make it possible to select the appropriate treatment.

The invention thus comprises a DNA chip according to the invention, characterized in that it contains, in addition, at least one nucleotide sequence of a microorganism different from Chlamydia trachomatis, immobilized on the support of the said chip; preferably, the different microorganism will be chosen from an associated microorganism, a bacterium of the Chlamydia family, and a variant of the species Chlamydia trachomatis.

Another subject of the present invention is a vector for the cloning and/or the expression of a sequence, characterized in that it contains a nucleotide sequence according to the invention.

Among the said vectors according to the invention, the vectors containing a nucleotide sequence encoding a polypeptide of the cellular, preferably outer, envelope of *Chlamydia trachomatis* or one of its representative fragments, are preferred.

In a specific embodiment, the vectors contain a nucleotide sequence encoding a Chlamydia trachomatis secreted polypeptide or one of its representative fragments or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its representative fragments, a polypeptide involved in lipopolysaccharide (LPS) biosynthesis, a Type III or non-Type III secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall anchored surface polypeptide, a polypeptide not found in Chlamydia pneumoniae, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids or of fatty acids of Chlamydia trachomatis or one of their representative fragments, or a polypeptide specific to Chlamydiae, are also preferred.

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According to the invention, the vectors comprise the elements necessary to allow the expression and/or the secretion of the said nucleotide sequences in a given host cell, and also form part of the invention.

The vector should, in this case, comprise a promoter, signals for initiation and for termination of translation, as well as appropriate regions for regulation of transcription. It should be capable of being stably maintained in the host cell and may optionally possess particular signals specifying the secretion of the translated protein. These different elements are chosen according to the host cell used. To this effect, the nucleotide sequences according to the invention may be inserted into autonomously-replicating vectors within the chosen host, or integrative vectors in the chosen host.

Any of the standard methods known to those skilled in the art for the insertion of DNA fragments into a vector may be used to construct expression vectors containing a chimeric gene consisting of appropriate transcriptional/translational control signals and the protein coding sequences. These methods may include in vitro recombinant DNA and synthetic techniques and in vivo recombinants (genetic recombination).

Expression of a polypeptide, peptide or derivative, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1 or ORFs contained within SEQ ID No. 1 may be regulated by a second nucleic acid sequence so that the protein or peptide is expressed in a host transformed with the recombinant DNA molecule. For example, expression of a protein or peptide may be controlled by any promoter/enhancer element known in the art. Promoters which may be used to control expression include, but are not limited to, the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 3' long terminal repeat of Rous sarcoma virus (Yamamoto, et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42); prokaryotic expression vectors such as the β-lactamase promoter (Villa-Kamaroff, et al., 1978, Proc. Natl. Acad. Sci. U.S.A. 75:3727-3731), or the tac promoter (DeBoer, et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:21-25); see also "Useful proteins from recombinant bacteria" in Scientific American, 1980, 242:74-94; plant expression vectors comprising the nopaline synthetase promoter region (Herrera-Estrella et al., 1983, Nature 303:209-213) or the cauliflower mosaic virus 35S RNA promoter (Gardner, et al., 1981, Nucl. 30 Acids Res. 2:2871), and the promoter of the photosynthetic enzyme ribulose biphosphate carboxylase (Herrera-Estrella et al., 1984, Nature 310:115-120); promoter elements from yeast or other fungi such as the Gal 4 promoter, the ADC (alcohol dehydrogenase) promoter, PGK (phosphoglycerol kinase) promoter, alkaline phosphatase promoter, and the following animal transcriptional control regions, which exhibit tissue specificity and have been utilized in transgenic animals: elastase I gene control 35 region which is active in pancreatic acinar cells (Swift et al., 1984, Cell 38:639-646; Ornitz et al.,

1986, Cold Spring Harbor Symp. Quant. Biol. 50:399-409; MacDonald, 1987, Hepatology 7:425-515); insulin gene control region which is active in pancreatic beta cells (Hanahan, 1985, Nature 315:115-122), immunoglobulin gene control region which is active in lymphoid cells (Grosschedl et al., 1984, Cell 38:647-658; Adames et al., 1985, Nature 318:533-538; Alexander et al., 1987, Mol. Cell. Biol. 7:1436-1444), mouse mammary tumor virus control region which is active in testicular, breast, lymphoid and mast cells (Leder et al., 1986, Cell 45:485-495), albumin gene control region which is active in liver (Pinkert et al., 1987, Genes and Devel. 1:268-276), alpha-fetoprotein gene control region which is active in liver (Krumlauf et al., 1985, Mol. Cell. Biol. 5:1639-1648; Hammer et al., 1987, Science 235:53-58; alpha 1-antitrypsin gene control region which is active in the liver 10 (Kelsey et al., 1987, Genes and Devel. 1:161-171), beta-globin gene control region which is active in myeloid cells (Mogram et al., 1985, Nature 315:338-340; Kollias et al., 1986, Cell 46:89-94; myelin basic protein gene control region which is active in oligodendrocyte cells in the brain (Readhead et al., 1987, Cell 48:703-712); myosin light chain-2 gene control region which is active in skeletal muscle (Sani, 1985, Nature 314:283-286), and gonadotropic releasing hormone gene control region 15 which is active in the hypothalamus (Mason et al., 1986, Science 234:1372-1378).

The vectors according to the invention are, for example, vectors of plasmid or viral origin. In a specific embodiment, a vector is used that comprises a promoter operably linked to a protein or peptide-encoding nucleic acid sequence in SEQ ID No. 1, or ORFs contained within SEQ ID No. 1, one or more origins of replication, and, optionally, one or more selectable markers (e.g., an antibiotic resistance gene). Expression vectors comprise regulatory sequences that control gene expression, including gene expression in a desired host cell. Preferred vectors for the expression of the polypeptides of the invention include the pET-type plasmid vectors (Promega) or pBAD plasmid vectors (Invitrogen). Furthermore, the vectors according to the invention are useful for transforming host cells so as to clone or express the nucleotide sequences of the invention.

Expression can also be achieved using targeted homologous recombination to activate Chlamydia trachomatis genes present in the cloned genomic DNA. A heterologous regulatory element may be inserted into a stable cell line or cloned microorganism, such that it is operatively linked with an endogenous Chlamydia trachomatis gene present in the cloned genome, using techniques, such as targeted homologous recombination, which are well known to those of skill in the 30 art (See, e.g., Chappel, U.S. Patent No. 4,215,051 and Skoultchi, WO 91/06667 each of which is incorporated herein in its entirety).

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Expression vector/host cell systems containing inserts of polynucleotide sequences in SEQ ID No. 1 or ORFs within SEQ ID No. 1, which encode polypeptides, peptides or derivatives, or analogs thereof, can be identified by three general approaches: (a) nucleic acid hybridization, (b) presence or absence of "marker" gene functions, and (c) expression of inserted sequences. In the first

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approach, the presence of a polynucleotide sequence inserted in an expression vector can be detected by nucleic acid hybridization using probes comprising sequences that are homologous to an inserted polynucleotide sequence. In the second approach, the recombinant vector/host system can be identified and selected based upon the presence or absence of certain "marker" gene functions (e.g., thymidine kinase activity, resistance to antibiotics, transformation phenotype, occlusion body formation in baculovirus, etc.) caused by the insertion of a polynucleotide sequence in the vector. For example, if the polynucleotide sequence in SEQ ID No. 1 or ORFs within SEQ ID No. 1 is inserted within the marker gene sequence of the vector, recombinants containing the insert can be identified by the absence of the marker gene function. In the third approach, recombinant expression vectors can be identified by assaying the product of the polynucleotide sequence expressed by the recombinant. Such assays can be based, for example, on the physical or functional properties of the expressed polypeptide in in vitro assay systems, e.g., binding with antibody, promotion of cell proliferation.

Once a particular recombinant DNA molecule is identified and isolated, several methods known in the art may be used to propagate it. The clones identified may be introduced into an appropriate host cell by standard methods, such as for example lipofection, electroporation, and heat shock. Once a suitable host system and growth conditions are established, recombinant expression vectors can be propagated and prepared in quantity.

The invention also encompasses the host cells transformed by a vector according to the invention. These cells may be obtained by introducing into host cells a nucleotide sequence inserted into a vector as defined above, and then culturing the said cells under conditions allowing the replication and/or the expression of the transfected nucleotide sequence.

The host cell may be chosen from eukaryotic or prokaryotic systems, such as for example bacterial cells (Olins and Lee, 1993), but also yeast cells (Buckholz, 1993), as well as animal cells, in particular cultures of mammalian cells (Edwards and Aruffo, 1993), and in particular Chinese hamster ovary (CHO) cells, but also insect cells in which methods using baculoviruses for example may be used (Luckow, 1993).

Furthermore, a host cell strain may be chosen which modulates the expression of the inserted sequences, or modifies and processes the gene product in the specific fashion desired. Expression from certain promoters can be elevated in the presence of certain inducers; thus, expression of the genetically engineered polypeptide may be controlled. Furthermore, different host cells have characteristic and specific mechanisms for the translational and post-translational processing and modification (e.g., glycosylation, phosphorylation) of proteins. Appropriate cell lines or host systems can be chosen to ensure the desired modification and processing of the foreign protein expressed. For example, expression in a bacterial system can be used to produce an unglycosylated core protein product. Expression in yeast will produce a glycosylated product. Expression in

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mammalian cells can be used to ensure "native" glycosylation of a heterologous protein. Furthermore, different vector/host expression systems may effect processing reactions to different extents.

A preferred host cell for the expression of the proteins of the invention consists of prokaryotic cells, such as Gram negative bacteria.

A further preferred host cell according to the invention is a bacterium belonging to the Chlamydia family, more preferably belonging to the species Chlamydia trachomatis or chosen from a microorganism associated with the species Chlamydia trachomatis.

In other specific embodiments, the polypeptides, peptides or derivatives, or analogs thereof may be expressed as a fusion, or chimeric protein product (comprising the protein, fragment, 10 analog, or derivative joined via a peptide bond to a heterologous protein sequence (of a different protein)). Such a chimeric product can be made by ligating the appropriate nucleic acid sequences encoding the desired amino acid sequences to each other by methods known in the art, in the proper coding frame, and expressing the chimeric product by methods commonly known in the art. Alternatively, such a chimeric product may be made by protein synthetic techniques, e.g., by use of a peptide synthesizer.

Genomic sequences can be cloned and expressed as translational gene products (i.e., peptides, polypeptides, and proteins) or transcriptional gene products (i.e., antisense and ribozymes).

The invention further relates to the intracellular production of an antisense nucleic acid sequence of SEQ ID No. 1 by transcription from an exogenous sequence. For example, a vector 20 can be introduced in vivo such that it is taken up by a cell, within which cell the vector or a portion thereof is transcribed, producing an antisense nucleic acid (RNA) of the invention. Such a vector would contain a sequence encoding an antisense nucleic acid. Such a vector can remain episomal or become chromosomally integrated, as long as it can be transcribed to produce the desired antisense RNA. Such vectors can be constructed by recombinant DNA technology methods standard in the art. Vectors can be plasmid, viral, or others known in the art, used for replication and expression in mammalian cells. Expression of the sequence encoding the antisense RNA can be by any promoter known in the art to act in mammalian, preferably human, cells. Such promoters can be inducible or constitutive. Such promoters include but are not limited to: the CMV promoter, the SV40 early promoter region (Bernoist and Chambon, 1981, Nature 290:304-310), the promoter contained in the 30 3N long terminal repeat of Rous sarcoma virus (Yamamoto et al., 1980, Cell 22:787-797), the herpes thymidine kinase promoter (Wagner et al., 1981, Proc. Natl. Acad. Sci. U.S.A. 78:1441-1445), the regulatory sequences of the metallothionein gene (Brinster et al., 1982, Nature 296:39-42), etc.

In a specific embodiment, the antisense oligonucleotide comprises catalytic RNA, or a ribozyme (see, e.g., PCT International Publication WO 90/11364, published October 4, 1990; Sarver et al., 1990, Science 247:1222-1225). In another embodiment, the oligonucleotide is a 2N-0-

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methylribonucleotide (Inoue et al., 1987, Nucl. Acids Res. 15:6131-6148), or a chimeric RNA-DNA analog (Inoue et al., 1987, FEBS Lett. 215:327-330).

In another embodiment, the antisense nucleic acids of the invention comprise a sequence complementary to at least a portion of an RNA transcript of a polynucleotide sequence in SEQ ID No. 1. However, absolute complementarity, although preferred, is not required. A sequence "complementary to at least a portion of an RNA," as referred to herein, means a sequence having sufficient complementarity to be able to hybridize with the RNA, forming a stable duplex; in the case of double-stranded antisense nucleic acid sequence, a single strand of the duplex DNA may thus be tested, or triplex formation may be assayed. The ability to hybridize will depend on both the degree of complementarity and the length of the antisense nucleic acid. Generally, the longer the hybridizing nucleic acid, the more base mismatches with an RNA transcribed from SEQ ID No. 1 may contain and still form a stable duplex (or triplex, as the case may be). One skilled in the art can ascertain a tolerable degree of mismatch by use of standard procedures to determine the melting point of the hybridized complex.

The invention also relates to the animals, except humans, comprising one of the above-described transformed cells according to the invention.

The production of transgenic animals according to the invention overexpressing one or more of the *Chlamydia trachomatis* genes will be preferably carried out on rats, mice or rabbits according to methods well known to persons skilled in the art such as viral or nonviral transfections.

The transgenic animals overexpressing one or more of the said genes may be obtained by transfection of multiple copies of the said genes under the control of a powerful promoter of a ubiquitous nature, or which is selective for one type of tissue. The transgenic animals may also be obtained by homologous recombination on embryonic stem cells, transfer of these stem cells to embryos, selection of the chimeras affected at the level of the reproductive lines, and growth of the said chimeras.

The transformed cells as well as the transgenic animals according to the invention can be used in methods of preparing the recombinant polypeptide.

It is now possible to produce recombinant polypeptides in a relatively large quantity by genetic engineering using the cells transformed with expression vectors according to the invention or using transgenic animals according to the invention.

The methods of preparing a polypeptide of the invention in recombinant form, characterized in that they use a vector and/or a cell transformed with a vector according to the invention and/or a transgenic animal comprising one of the said transformed cells according to the invention, are themselves included in the present invention.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or

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a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a polypeptide of the cellular envelope of Chlamydia trachomatis or one of its representative fragments, more preferably encoding a polypeptide of the outer cellular envelope of Chlamydia trachomatis or one of its fragment, are preferred.

Among the said methods of preparing a polypeptide of the invention in recombinant form, the methods of preparation using a vector, and/or a cell transformed with the said vector and/or a transgenic animal comprising one of the said transformed cells, containing a nucleotide sequence encoding a Chlamydia trachomatis secreted polypeptide or one of its representative fragments, or encoding a transport polypeptide, a surface exposed polypeptide, a lipoprotein or one of its representative fragments, a polypeptide involved in lipopolysaccharide biosynthesis, a Type III or 10 other secreted polypeptide, a polypeptide containing RGD attachment sites, a cell wall anchored surface polypeptide, a polypeptide not found in Chlamydia pneumoniae, a ribosomal polypeptide or a polypeptide involved in secretion, transcription, translation, maturation of proteins, a polypeptide involved in the synthesis of the wall, a polypeptide involved in the virulence, a polypeptide involved 15 in the intermediate metabolism, in particular in the metabolism of sugars and/or of cofactors, a polypeptide involved in the metabolism of nucleotides, of amino acids, of nucleic acids or of fatty acids of Chlamydia trachomatis or one of their representative fragments, or a polypeptide specific to Chlamydiae, are also preferred.

The recombinant polypeptides obtained as indicated above may be provided either in glycosylated or nonglycosylated form and may or may not have the natural tertiary structure.

A preferred variant consists in producing a recombinant polypeptide fused to a «carrier» protein (chimeric protein). The advantage of this system is that it allows a stabilization and a reduction in proteolysis of the recombinant product, an increase in solubility during renaturation in vitro and/or a simplification of purification when the fusion partner has affinity for a specific ligand.

More particularly, the invention relates to a method of preparing a polypeptide of the invention comprising the following steps:

- a) culture of the transformed cells under conditions allowing the expression of a recombinant polypeptide having a nucleic acid sequence according to the invention;
- b) where appropriate, recovery of the said recombinant polypeptide.

When the method of preparing a polypeptide of the invention uses a transgenic animal according to the invention, the recombinant polypeptide is then extracted from the said animal.

The subject of the invention is also a polypeptide capable of being obtained by a method of the invention as described above.

35 The invention also comprises a method of preparing a synthetic polypeptide,

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characterized in that it uses an amino acid sequence of polypeptides according to the invention.

The invention also relates to a synthetic polypeptide obtained by a method according to the invention.

Polypeptides according to the invention may also be prepared by conventional 5 techniques in the field of peptide synthesis under conditions suitable to produce the polypeptides encoded by the polynucleotide of the invention. This synthesis may be carried out in and recovered from a homogeneous solution or on a solid phase.

For example, the synthesis technique in a homogeneous solution described by Houbenweyl in 1974 may be used.

This method of synthesis consists in successively condensing, in pairs, the successive amino acids in the required order, or in condensing amino acids and fragments previously formed and already containing several amino acids in the appropriate order, or alternatively several fragments thus previously prepared, it being understood that care will have been taken to protect beforehand all the reactive functional groups carried by these amino acids or fragments, with the exception of the amine functional groups of one and the carboxyl functional groups of the other or vice versa, which should normally take part in the formation of the peptide bonds, in particular after activation of the carboxyl functional group, according to methods well known in peptide synthesis.

According to another preferred technique of the invention, the one described by Merrifield is used.

To manufacture a peptide chain according to the Merrifield method, a highly porous polymer resin is used, onto which the first C-terminal amino acid of the chain is attached. This amino acid is attached onto a resin via its carboxyl group and its amine functional group is protected. The amino acids which will constitute the peptide chain are thus attached, one after another, onto the amine group, each time deprotected beforehand, of the portion of the peptide chain already formed, 25 and which is attached to the resin. When the entire peptide chain desired is formed, the protecting groups are removed from the various amino acids constituting the peptide chain and the peptide is detached from the resin with the aid of an acid.

The invention relates, in addition, to hybrid (fusion) polypeptides having at least one polypeptide or one of its representative fragments according to the invention, and a sequence of a 30 polypeptide capable of eliciting an immune response in humans or animals.

Advantageously, the antigenic determinant is such that it is capable of eliciting a humoral and/or cellular response.

An antigenic determinant may be identified by screening expression libraries of the Chlamydia trachomatis genome with antibodies contained in the serum of patients infected with a bacterium belonging to the species Chlamydia trachomatis. An antigenic determinant may comprise a WO 99/28475 PCT/IB98/01939

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polypeptide or one of its fragments according to the invention, in glycosylated form, used in order to obtain immunogenic compositions capable of inducing the synthesis of antibodies directed against multiple epitopes. The said polypeptides or their glycosylated fragments also form part of the invention.

These hybrid molecules may consist, in part, of a carrier molecule for polypeptides or for their representative fragments according to the invention, combined with a portion which may be immunogenic, in particular an epitope of the diphtheria toxin, the tetanus toxin, a hepatitis B virus surface antigen (patent FR 79 21811), the poliomyelitis virus VP1 antigen or any other viral or bacterial toxin or antigen.

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The methods of synthesizing the hybrid molecules include the methods used in genetic engineering to construct hybrid nucleotide sequences encoding the desired polypeptide sequences. Reference may be advantageously made, for example, to the technique for producing genes encoding fusion proteins described by Minton in 1984.

The said hybrid nucleotide sequences encoding a hybrid polypeptide as well as the hybrid polypeptides according to the invention, characterized in that they are recombinant polypeptides obtained by the expression of the said hybrid nucleotide sequences, also form part of the invention.

The invention also comprises the vectors characterized in that they contain one of the said hybrid nucleotide sequences. The host cells transformed by the said vectors, the transgenic animals comprising one of the said transformed cells as well as the methods of preparing recombinant polypeptides using the said vectors, the said transformed cells and/or the said transgenic animals of course also form part of the invention.

The polypeptides according to the invention, the antibodies according to the invention described below and the nucleotide sequences according to the invention may advantageously be used in *in vitro* and/or *in vivo* methods for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis*, in a biological sample (biological tissue or fluid) which is likely to contain them. These methods, depending on the specificity of the polypeptides, of the antibodies and of the nucleotide sequences according to the invention which will be used, may in particular detect and/or identify the bacterial variants belonging to the species *Chlamydia trachomatis* as well as the associated microorganisms capable of being detected by the polypeptides, the antibodies and the nucleotide sequences according to the invention which will be chosen. It may, for example, be advantageous to choose a polypeptide, an antibody or a nucleotide sequence according to the invention, which is capable of detecting any bacterium of the *Chlamydia* family by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the family or, on the contrary, it will be most particularly advantageous to target a variant of the

species Chlamydia trachomatis, which is responsible, for example, for the induction or the worsening of pathologies specific to the targeted variant, by choosing a polypeptide, an antibody and/or a nucleotide sequence according to the invention which is specific to the said variant.

The polypeptides according to the invention may advantageously be used in a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, in a biological sample (biological tissue or fluid) which is likely to contain them, characterized in that it comprises the following steps:

- a) bringing this biological sample into contact with a polypeptide or one of its representative fragments according to the invention (under conditions allowing an immunological reaction between the said polypeptide and the antibodies which may be present in the biological sample);
- b) detecting the antigen-antibody complexes which may be formed.

Preferably, the biological sample consists of a fluid, for example a human or animal serum, blood or biopsies.

Any conventional procedure may be used to carry out such a detection of the antigenantibody complexes which may be formed.

By way of example, a preferred method uses immunoenzymatic procedures based on the ELISA technique, immunofluorescence procedures or radioimmunological procedures (RIA), and the like.

Accordingly, the invention also relates to the polypeptides according to the invention,
labelled with the aid of a suitable label such as a label of the enzymatic, fluorescent or radioactive
type.

Such methods comprise, for example, the following steps:

- deposition of defined quantities of a polypeptide composition according to the invention into the wells of a microtitre plate,
- 25 introduction, into the said wells, of increasing dilutions of serum, or of a different biological sample as defined above, which has to be analysed,
  - incubation of the microplate,
  - introduction, into the wells of the microtitre plate, of labelled antibodies directed against human or animal immunoglobulins, these antibodies having been labelled with the aid of an enzyme selected from those which are capable of hydrolyzing a substrate, thereby modifying the absorption of the radiation of the latter, at least at a defined wavelength, for example at 550 nm,
    - detection, by comparison with a control, of the quantity of substrate hydrolyzed.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a polypeptide according to the invention,
- where appropriate, the reagents for constituting the medium appropriate for the immunological or specific reaction,
- the reagents allowing the detection of the antigen-antibody complexes produced by the immunological reaction between the polypeptide(s) of the invention and the antibodies which may be present in the biological sample, it being possible for these reagents also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the polypeptide according to the invention is not labelled,
- where appropriate, a reference biological sample (negative control) free of antibodies recognized by a polypeptide according to the invention,
  - where appropriate, a reference biological sample (positive control) containing a predetermined quantity of antibodies recognized by a polypeptide according to the invention.

According to the invention, the polypeptides, peptides, fusion proteins or other derivatives, or analogs thereof encoded by a polynucleotide sequence in SEQ ID No. 1, may be used as an immunogen to generate antibodies which immunospecifically bind such an immunogen. Such antibodies may include, but are not limited to, polyclonal and monoclonal antibodies, humanized or chimeric antibodies, single chain antibodies, Fab fragments, F(ab)<sub>2</sub> fragments, fragments produced by a Fab expression library, anti-idiotypic (anti-Id) antibodies, and epitope-binding fragments of any of the above. In a specific embodiment, the antibody to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1 is a bispecific antibody (see generally, e.g. Fanger and Drakeman, 1995, Drug News and Perspectives 8: 133-137). Such a bispecific antibody is genetically engineered to recognize both (1) an epitope and (2) one of a variety of "trigger" molecules, e.g. Fc receptors on myeloid cells, and CD3 and CD2 on T cells, that have been identified as being able to cause a cytotoxic T-cell to destroy a particular target. Such bispecific antibodies can be prepared either by chemical conjugation, hybridoma, or recombinant molecular biology techniques known to the skilled artisan.

Various procedures known in the art may be used for the production of polyclonal antibodies to a polypeptide, peptide or other derivative, or analog thereof encoded by a polynucleotide sequence in SEQ ID No. 1. For the production of antibody, various host animals can be immunized by injection with a polypeptide, or peptide or other derivative, or analog thereof, including but not limited to rabbits, mice, rats, etc. Various adjuvants, depending on the host species, may be used to increase the immunological response, including but not limited to Stimulon QS-21 (Aquila Biopharmaceuticals, Inc., Framingham, MA), MPL (3-O-deacylated monophosphoryl lipid A; RIBI ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA), Freund's (complete and incomplete), mineral gels such as aluminum hydroxide,

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surface active substances such as lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, keyhole limpet hemocyanins, dinitrophenol, BCG (bacille Calmette-Guerin), and corynebacterium parvum. Alternatively, polyclonal antibodies may be prepared by purifying, on an affinity column onto which a polypeptide according to the invention has been previously attached, the antibodies contained in the serum of patients infected with a bacterium belonging to the species *Chlamydia trachomatis*.

For preparation of monoclonal antibodies directed toward a polypeptide, peptide or other derivative, or analog, any technique which provides for the production of antibody molecules by continuous cell lines in culture may be used. For example, the hybridoma technique originally developed by Kohler and Milstein (1975, Nature 256:495-497), as well as the trioma technique, the human B-cell hybridoma technique (Kozbor et al., 1983, Immunology Today 4:72), and the EBVhybridoma technique to produce human monoclonal antibodies (Cole et al., 1985, in Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, Inc., pp. 77-96). In an additional embodiment of the invention, monoclonal antibodies can be produced in germ-free animals utilizing technology 15 described in PCT/US90/02545. In another embodiment of the invention, transgenic non-human animals can be used for the production of human antibodies utilizing technology described in WO 98/24893 and WO 96/33735. According to the invention, human antibodies may be used and can be obtained by using human hybridomas (Cote et al., 1983, Proc. Natl. Acad. Sci. U.S.A. 80:2026-2030) or by transforming human B cells with EBV virus in vitro (Cole et al., 1985, in Monoclonal Antibodies and Cancer Therapy, Alan R. Liss, pp. 77-96). In fact, according to the invention, techniques developed for the production of «chimeric antibodies», (Morrison et al., 1984, PROC. NATL. ACAD. SCI. U.S.A. 81:6851-6855; Neuberger et al., 1984, Nature 312:604-608; Takeda et al., 1985, Nature 314:452-454) by splicing the genes from a mouse antibody molecule specific for a polypeptide, peptide or other derivative, or analog together with genes from a human antibody molecule of appropriate biological activity can be used; such antibodies are within the scope of this invention.

According to the invention, techniques described for the production of single chain antibodies (U.S. Patent 4,946,778) can be adapted to produce polypeptide or peptide-specific single chain antibodies. An additional embodiment of the invention utilizes the techniques described for the construction of Fab expression libraries (Huse et al., 1989, Science 246:1275-1281) to allow rapid and easy identification of monoclonal Fab fragments with the desired specificity for polypeptides, derivatives, or analogs.

Antibody fragments which contain the idiotype of the molecule can be generated by known techniques. For example, such fragments include but are not limited to: the F(ab')<sub>2</sub> fragment which can be produced by pepsin digestion of the antibody molecule; the Fab' fragments which can be

generated by reducing the disulfide bridges of the F(ab')<sub>2</sub> fragment, the Fab fragments which can be generated by treating the antibody molecule with papain and a reducing agent, and Fv fragments.

In addition, techniques have been developed for the production of chimerized (See, e.g., Boss, M. et al., U.S. Patent No. 4,816,397; and Cabilly, S. et al., U.S. Patent No. 5,585,089 each of which is incorporated herein by reference in its entirety) humanized antibodies (See, e.g., Queen, U.S. Patent No. 5,585,089, which is incorporated herein by reference in its entirety.) An immunoglobulin light or heavy chain variable region consists of a "framework" region interrupted by three hypervariable regions, referred to as complementarily determining regions (CDRs). The extent of the framework region and CDRs have been precisely defined (See, "Sequences of Proteins of Immunological Interest", Kabat, E. et al., U.S. Department of Health and Human Services (1983)). Briefly, humanized antibodies are antibody molecules from non-human species having one or more CDRs from the non-human species and a framework from a human immunoglobulin molecule.

The antibodies of the invention may also be labelled in the same manner as described above for the nucleic probes of the invention such as an enzymatic, fluorescent or radioactive type labelling.

The invention relates, in addition, to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:

- a) bringing the biological sample (biological tissue or fluid) into contact with a mono- or polyclonal antibody according to the invention (under conditions allowing an immunological reaction between the said antibodies and the polypeptides of the bacterium belonging to the species *Chlamydia trachomatis* or to an associated microorganism which may be present in the biological sample, that is, under conditions suitable for the formation of immune complexes);
- b) detecting the antigen-antibody complex which may be formed.
- Also falling within the scope of the invention is a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:
  - a polyclonal or monoclonal antibody according to the invention, labelled where appropriate;
- where appropriate, a reagent for constituting the medium appropriate for carrying out the 30 immunological reaction;
  - a reagent allowing the detection of the antigen-antibody complexes produced by the immunological reaction, it being possible for this reagent also to carry a label, or to be capable of being recognized in turn by a labelled reagent, more particularly in the case where the said monoclonal or polyclonal antibody is not labelled;
- 35 where appropriate, reagents for carrying out the lysis of the cells in the sample tested.

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The principle of the DNA chip which was explained above may also be used to produce protein "chips" on which the support has been coated with a polypeptide or an antibody according to the invention, or arrays thereof, in place of the DNA. These protein "chips" make it possible, for example, to analyse the biomolecular interactions (BIA) induced by the affinity capture of target analytes onto a support coated, for example, with proteins, by surface plasma resonance (SPR). Reference may be made, for example, to the techniques for coupling proteins onto a solid support which are described in EP 524 800 or to the methods describing the use of biosensor-type protein chips such as the BIAcore-type technique (Pharmacia) (Arlinghaus et al., 1997, Krone et al., 1997, Chatelier et al., 1995). These polypeptides or antibodies according to the invention, capable of specifically binding antibodies or polypeptides derived from the sample to be analysed, may thus be used in protein chips for the detection and/or the identification of proteins in samples. The said protein chips may in particular be used for infectious diagnosis and may preferably contain, per chip, several polypeptides and/or antibodies of the invention of different specificity, and/or polypeptides and/or antibodies capable of recognizing microorganisms different from Chlamydia trachomatis.

Accordingly, the subject of the present invention is also the polypeptides and the antibodies according to the invention, characterized in that they are immobilized on a support, in particular of a protein chip.

The protein chips, characterized in that they contain at least one polypeptide or one antibody according to the invention immobilized on the support of the said chip, also form part of the invention.

The invention comprises, in addition, a protein chip according to the invention, characterized in that it contains, in addition, at least one polypeptide of a microorganism different from *Chlamydia trachomatis* or at least one antibody directed against a compound of a microorganism different from *Chlamydia trachomatis*, immobilized on the support of the said chip.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a protein chip according to the invention.

The subject of the present invention is also a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it uses a nucleotide sequence according to the invention.

More particularly, the invention relates to a method for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism in a biological sample, characterized in that it comprises the following steps:

- a) where appropriate, isolation of the DNA from the biological sample to be analysed, or optionally production of a cDNA from the RNA in the biological sample;
- b) specific amplification of the DNA of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism with the aid of at least one primer according to the invention;
- 5 c) detection of the amplification products.

These may be detected, for example, by the molecular hybridization technique using a nucleic probe according to the invention. This probe will be advantageously labelled with a nonradioactive (cold probe) or radioactive element.

For the purposes of the present invention, "DNA in the biological sample" or "DNA contained in the biological sample" will be understood to mean either the DNA present in the biological sample considered, or optionally the cDNA obtained after the action of a reverse transcriptase-type enzyme on the RNA present in the said biological sample.

Another aim of the present invention consists in a method according to the invention, characterized in that it comprises the following steps:

- 15 a) bringing a nucleotide probe according to the invention into contact with a biological sample, the DNA contained in the biological sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to complementary base pairs of the DNA of a bacterium belonging to the species *Chlamydia trachomatis* or to an associated microorganism;
- 20 b) detecting the hybridization complex formed between the nucleotide probe and the DNA in the biological sample.

The present invention also relates to a method according to the invention, characterized in that it comprises the following steps:

- a) bringing a nucleotide probe immobilized on a support according to the invention into contact
   with a biological sample, the DNA in the sample having, where appropriate, been previously made accessible to hybridization, under conditions allowing the hybridization of the probe to the DNA of a bacterium belonging to the species Chlamydia trachomatis or to an associated microorganism;
  - b) bringing the hybrid formed between the nucleotide probe immobilized on a support and the DNA contained in the biological sample, where appropriate after removal of the DNA in the biological sample which has not hybridized with the probe, into contact with a labelled nucleotide probe according to the invention;
  - c) detecting the new hybrid formed in step b).

According to an advantageous embodiment of the method for the detection and/or the identification defined above, it is characterized in that, prior to step a), the DNA in the biological sample is primer-extended and/or amplified beforehand with the aid of at least one primer according

to the invention.

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The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- 5 a) a nucleotide probe according to the invention;
  - b) where appropriate, the reagents necessary for carrying out a hybridization reaction;
  - c) where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a) a nucleotide probe, called capture probe, according to the invention:
- b) an oligonucleotide probe, called detection probe, according to the invention;
- c) where appropriate, at least one primer according to the invention as well as the reagents (e.g., polymerase and/or deoxynucleotide triphosphates) necessary for a DNA amplification reaction.

The invention also relates to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, characterized in that it comprises the following components:

- a) at least one primer according to the invention;
- 20 b) where appropriate, the reagents necessary for carrying out a DNA amplification reaction;
  - c) where appropriate, a component which makes it possible to check the sequence of the amplified fragment, more particularly an oligonucleotide probe according to the invention.

The invention relates, in addition, to a kit or set for the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* or to an associated microorganism, or for the detection and/or the identification of a microorganism characterized in that it comprises a DNA chip according to the invention.

The invention also relates to a method or to a kit or set according to the invention for the detection and/or the identification of bacteria belonging to the species Chlamydia trachomatis, characterized in that the said primer and/or the said probe according to the invention are chosen from the nucleotide sequences specific to the species Chlamydia trachomatis, in that the said polypeptides according to the invention are chosen from the polypeptides specific to the species Chlamydia trachomatis and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides specific to the species Chlamydia trachomatis.

Preferably, the said method or the said kit or set above according to the invention, for

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the detection and/or the identification of bacteria belonging to the species *Chlamydia trachomatis* is characterized in that the said primer and/or the said probe or the said polypeptides are chosen from the nucleotide sequences or polypeptides according to the invention which have been identified as being specific to the species *Chlamydia trachomatis* and in that the said antibodies according to the invention are chosen from the antibodies directed against the polypeptides according to the invention chosen from the polypeptides identified as being specific to the species *Chlamydia trachomatis*.

The invention relates, in addition, to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of a condition caused by, genital diseases which are induced or worsened by a *Chlamydia trachomatis* infection.

The invention also relates to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of conditions caused by, eye diseases induced or worsened by a *Chlamydia trachomatis* infection.

The invention also relates to a method or a kit or set according to the invention for the diagnosis of predispositions to, or of conditions caused by, systemic diseases, in particular of the lymphatic system, which are induced or worsened by a *Chlamydia trachomatis* infection.

According to another aspect, the subject of the invention is the use of polypeptides according to the invention, of cells transformed with a vector according to the invention and/or of transformed animals according to the invention, for the biosynthesis or the biodegradation of organic or inorganic compounds.

As has been mentioned above, the nucleotide sequences of the invention were identified by homology with sequences known to encode, for example, polypeptides or fragments of enzymatic polypeptides involved in the biosynthesis or the biodegradation of organic or inorganic molecules.

It is thus possible to use the said polypeptides of the invention in a similar manner for the biosynthesis or the biodegradation of organic or inorganic compounds of industrial or therapeutic interest (called compounds of interest).

Among these polypeptides, there may be mentioned in particular the enzymes involved in metabolism, such as the proteolytic enzymes, amino transferases, glucose metabolism, or the enzymes which may be used in the biosynthesis of sugars, amino acids, fatty acids, polypeptides, nucleotides, nucleic acids or any other organic or inorganic compound or in the biodegradation of organic or inorganic compounds.

Among these polypeptides, there may be mentioned, in addition, the mutated or modified enzymes corresponding to mutated or modified polypeptides according to the invention which may also be used for the biosynthesis or the biodegradation of organic or inorganic compounds at the industrial level, such as, for example, the production of compounds of interest, the reprocessing

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of manufacturing residues applied to the food industries, to the papermaking industry or to the chemical and pharmaceutical industries.

The methods of biosynthesis or biodegradation of organic or inorganic compounds, characterized in that they use a polypeptide or one of its representative fragments according to the invention, transformed cells according to the invention and/or a transformed animal according to the invention, also form part of the invention.

The invention relates, in addition, to the use of a nucleotide sequence according to the invention, of a polypeptide according to the invention, of an antibody according to the invention, of a cell according to the invention, and/or of a transformed animal according to the invention, for the selection of an organic or inorganic compound capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the cellular replication of eukaryotic or prokaryotic cells or capable of inducing, inhibiting or worsening the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms.

The invention also comprises screening assays that comprise method of selecting compounds capable of binding to a polypeptide, fusion polypeptide, or one of its representative fragments according to the invention, capable of binding to a nucleotide sequence according to the invention, or capable of recognizing an antibody according to the invention, and/or capable of modulating, regulating, inducing or inhibiting the expression of genes, and/or of modifying the growth or the cellular replication of eukaryotic or prokaryotic cells, or capable of inducing, inhibiting or worsening, in an animal or human organism, the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms, characterized in that it comprises the following steps:

- a) bringing the said compound into contact with the said polypeptide, the said nucleotide sequence, with a transformed cell according to the invention and/or administering the said compound to a transformed animal according to the invention;
- b) determining the capacity of the said compound to bind with the said polypeptide or the said nucleotide sequence, or to modulate, regulate, induce or inhibit the expression of genes, or to modulate growth or cellular replication, or to induce, inhibit or worsen in the said transformed animal, the pathologies linked to an infection by *Chlamydia trachomatis* or one of its associated microorganisms.

The transformed cells and/or animals according to the invention may advantageously serve as a model and may be used in methods for studying, identifying and/or selecting compounds capable of being responsible for pathologies induced or worsened by *Chlamydia trachomatis*, or capable of preventing and/or of treating these pathologies such as, for example, genital, eye or systemic diseases, especially of the lymphatic system. In particular, the transformed host cells, in

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particular bacteria of the *Chlamydia* family whose transformation with a vector according to the invention may, for example, increase or inhibit its infectivity, or modulate the pathologies usually induced or worsened by the infection, may be used to infect animals in which the onset of pathologies will be monitored. These nontransformed animals, infected for example with transformed *Chlamydia* bacteria, may serve as a study model. In the same manner, the transformed animals according to the invention may, for example, exhibit predispositions to genital and/or eye and/or systemic diseases, especially of the lymphatic system, and thus be used in methods for selecting compounds capable of preventing and/or of treating the said diseases. The said methods using the said transformed cells and/or transformed animals form part of the invention.

The compounds capable of being selected may be organic compounds such as polypeptides or carbohydrates or any other organic or inorganic compounds already known, or new organic compounds produced using molecular modelling techniques and obtained by chemical or biochemical synthesis, these techniques being known to persons skilled in the art.

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The said selected compounds may be used to modulate the growth and/or the cellular replication of *Chlamydia trachomatis* or any other associated microorganism and thus to control infection by these microorganisms. The said compounds according to the invention may also be used to modulate the growth and/or the cellular replication of all eukaryotic or prokaryotic cells, in particular tumour cells and infectious microorganisms, for which the said compounds will prove active, the methods which make it possible to determine the said modulations being well known to persons skilled in the art.

Compound capable of modulating the growth of a microorganism is understood to designate any compound which makes it possible to act, to modify, to limit and/or to reduce the development, the growth, the rate of proliferation and/or the viability of the said microorganism.

This modulation may be achieved, for example, by an agent capable of binding to a protein and thus of inhibiting or of potentiating its biological activity, or capable of binding to a membrane protein of the outer surface of a microorganism and of blocking the penetration of the said microorganism into the host cell or of promoting the action of the immune system of the infected organism directed against the said microorganism. This modulation may also be achieved by an agent capable of binding to a nucleotide sequence of a DNA or RNA of a microorganism and of blocking, for example, the expression of a polypeptide whose biological or structural activity is necessary for the growth or for the reproduction of the said microorganism.

Associated microorganism is understood to designate in the present invention any microorganism whose gene expression may be modulated, regulated, induced or inhibited, or whose growth or cellular replication may also be modulated by a compound of the invention. Associated microorganism is also understood to designate in the present invention any microorganism containing

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nucleotide sequences or polypeptides according to the invention. These microorganisms may, in some cases, contain polypeptides or nucleotide sequences identical or homologous to those of the invention may also be detected and/or identified by the detection and/or identification methods or kit according to the invention and may also serve as a target for the compounds of the invention.

The invention relates to the compounds capable of being selected by a method of selection according to the invention.

The invention also relates to a pharmaceutical composition comprising a compound chosen from the following compounds:

a nucleotide sequence according to the invention;

10 a polypeptide or fusion polypeptide according to the invention;

a vector according to the invention;

an antibody according to the invention; and

a compound capable of being selected by a method of selection according to the invention, optionally in combination with a pharmaceutically acceptable vehicle or carrier.

An effective quantity is understood to designate a sufficient quantity of the said compound or antibody, or of a polypeptide of the invention, which makes it possible to modulate the growth of *Chlamydia trachomatis* or of an associated microorganism.

The invention also relates to a pharmaceutical composition according to the invention for the prevention or the treatment of an infection by a bacterium belonging to the species *Chlamydia trachomatis* or by an associated microorganism.

The invention relates, in addition, to an immunogenic and/or vaccine composition, characterized in that it comprises one or more polypeptides according to the invention and/or one or more hybrid polypeptides according to the invention.

The invention also comprises the use of a transformed cell according to the invention, for the preparation of a vaccine composition.

The invention also relates to a vaccine composition, characterized in that it contains a nucleotide sequence according to the invention, a vector according to the invention and/or a transformed cell according to the invention.

The invention also relates to the vaccine compositions according to the invention, for the prevention or the treatment of an infection by a bacterium belonging to the species *Chlamydia trachomatis* or by an associated microorganism.

The invention also relates to the use of DNA encoding polypeptides of *Chlamydia trachomatis*, in particular antigenic determinants, to be formulated as vaccine compositions. In accordance with this aspect of the invention, the DNA of interest is engineered into an expression vector under the control of regulatory elements, which will promote expression of the DNA, *i.e.*,

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promoter or enhancer elements. In one preferred embodiment, the promoter element may be cellspecific and permit substantial transcription of the DNA only in predetermined cells. The DNA may be introduced directly into the host either as naked DNA (U.S. Patent No. 5,679,647 incorporated herein by reference in their entirety) or formulated in compositions with other agents which may facilitate uptake of the DNA including viral vectors, i.e., adenovirus vectors, or agents which facilitate immunization, such as bupivicaine and other local anesthetics (U.S. Patent 5,593,972 incorporated herein by reference in their entirety), saponins (U.S. Patent 5,739,118 incorporated herein by reference in their entirety) and cationic polyamines (published international application WO 96/10038 incorporated herein by reference in their entirety).

The DNA sequence encoding the antigenic polypeptide and regulatory element may be inserted into a stable cell line or cloned microorganism, using techniques, such as targeted homologous recombination, which are well known to those of skill in the art, and described e.g., in Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

Such cell lines and microorganisms may be formulated for vaccine purposes. In yet another embodiment, the DNA sequence encoding the antigenic polypeptide and regulatory element may be delivered to a mammalian host and introduced into the host genome via homologous recombination (See, Chappel, U.S. Patent No. 4,215,051; Skoultchi, WO 91/06667 each of which is incorporated herein by reference in its entirety.

Preferably, the immunogenic and/or vaccine compositions according to the invention intended for the prevention and/or the treatment of an infection by Chlamydia trachomatis or by an associated microorganism will be chosen from the immunogenic and/or vaccine compositions comprising a polypeptide or one of its representative fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia trachomatis. The vaccine 25 compositions comprising nucleotide sequences will also preferably comprise nucleotide sequences encoding a polypeptide or one of its fragments corresponding to a protein, or one of its representative fragments, of the cellular envelope of Chlamydia trachomatis.

Among these preferred immunogenic and/or vaccine compositions, the most preferred are those comprising a polypeptide or one of its representative fragments, or a nucleotide sequence or 30 one of its representative fragments whose sequences are chosen from the nucleotide or amino acid sequences identified in this functional group and listed above.

The polypeptides of the invention or their representative fragments entering into the immunogenic compositions according to the invention may be selected by techniques known to persons skilled in the art, such as for example on the capacity of the said polypeptides to stimulate T 35 cells, which results, for example, in their proliferation or the secretion of interleukins, and which leads to the production of antibodies directed against the said polypeptides.

In mice, in which a weight dose of the vaccine composition comparable to the dose used in humans is administered, the antibody reaction is tested by collecting serum followed by a study of the formation of a complex between the antibodies present in the serum and the antigen of the vaccine composition, according to the customary techniques.

According to the invention, the said vaccine compositions will be preferably in combination with a pharmaceutically acceptable vehicle and, where appropriate, with one or more appropriate immunity adjuvants.

Various types of vaccines are currently available for protecting humans against infectious diseases: attenuated live microorganisms (M. bovis - BCG for tuberculosis), inactivated microorganisms (influenza virus), acellular extracts (Bordetella pertussis for whooping cough), recombinant proteins (hepatitis B virus surface antigen), polysaccharides (pneumococci). Experiments are underway on vaccines prepared from synthetic peptides or from genetically modified microorganisms expressing heterologous antigens. Even more recently, recombinant plasmid DNAs carrying genes encoding protective antigens were proposed as an alternative vaccine strategy. This type of vaccination is carried out with a particular plasmid derived from an E. coli plasmid which does not replicate in vivo and which encodes only the vaccinal protein. Animals were immunized by simply injecting the naked plasmid DNA into the muscle. This technique leads to the expression of the vaccine protein in situ and to a cell-type (CTL) and a humoral type (antibody) immune response.

This double induction of the immune response is one of the main advantages of the technique of vaccination with naked DNA.

The vaccine compositions of the present invention can be evaluated in in vitro and in vivo animal models prior to host, e.g., human, administration. For example, in vitro neutralization assays such as those described by Peterson et al. (1988) can be utilized. The assay described by Peterson et al. (1988) is suitable for testing vaccine compositions directed toward either Chlamydia trachomatis or Chlamydia pneumoniae.

Briefly, hyper-immune antisera is diluted in PBS containing 5% guinea pig serum, as a complement source. Chlamydiae (10<sup>4</sup> IFU; inclusion forming units) are added to the antisera dilutions. The antigen-antibody mixtures are incubated at 37°C for 45 minutes and inoculated into duplicate confluent Hep-2 or HeLa cell monolayers contained in glass vials (e.g., 15 by 45 mm), which have been washed twice with PBS prior to inoculation. The monolayer cells are infected by centrifugation at 1000X g for 1 hour followed by stationary incubation at 37° for 1 hour. Infected monolayers are incubated for 48 or 72 hours, fixed and stained with a Chlamydiae specific antibody, such as anti-MOMP for C. trachomatis, etc. Inclusion-bearing cells are counted in ten fields at a magnification of 200X. Neutralization titer is assigned based on the dilution that gives 50%

inhibition as compared to control monolayers/IFU.

The efficacy of vaccine compositions can be determined *in vivo* by challenging animal models of *Chlamydia trachomatis* infection, *e.g.*, guinea pigs or mice, with the vaccine compositions. For example, *in vivo* vaccine composition challenge studies in the guinea pig model of *Chlamydia trachomatis* infection can be performed. Briefly, female guinea pigs weighing 450 to 500 g are housed in an enviornmentally controlled room with a 12 hour light-dark cycle and immunized with vaccine compositions via a variety of immunization routes. Post-vaccination, guinea pigs are infected in the genital tract with the agent of guinea pig inclusion conjunctivitis (GPIC), which has been grown in HeLa or McCoy cells (Rank et al. (1988)). Each animal receives approximately 1.4x10<sup>7</sup> inclusion forming units (IFU) contained in 0.05 ml of sucrose-phosphate-glutamate buffer, pH 7.4 (Schacter, J. (1980)). The course of infection monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorescence with GPIC specific antisera, or by Giemsastained smear from a scraping from the genital tract (Rank et al. (1988)). Antibody titers in the serum is determined by an enzyme-linked immunosorbent assay.

Alternatively, in vivo vaccine composition challenge studies can be performed in the murine model of Chlamydia trachomatis (Morrison et al., 1995). Briefly, female mice 7 to 12 weeks of age receive 2.5 mg of depoprovera subcutaneously at 10 and 3 days before vaginal infection. Post-vaccination, mice are infected in the genital tract with 1,500 inclusion-forming units of Chlamydia trachomatis contained in 5ml of sucrose-phosphate-glutamate buffer, pH. 7.4. The course of infection is monitored by determining the percentage of inclusion-bearing cells by indirect immunofluorence with Chlamydia trachomatis specific antisera, or by a Giemsa-stained smear from a scraping from the genital tract of an infected mouse. The presence of antibody titers in the serum of a mouse is determined by an enzyme-linked immunosorbent assay.

The vaccine compositions comprising nucleotide sequences or vectors into which the said sequences are inserted are in particular described in International Application No. WO 90/11092 and also in International Application No. WO 95/11307.

The nucleotide sequence constituting the vaccine composition according to the invention may be injected into the host after having been coupled to compounds which promote the penetration of this polynucleotide inside the cell or its transport up to the cell nucleus. The resulting conjugates may be encapsulated into polymeric microparticles, as described in International Application No. WO 94/27238 (Medisorb Technologies International).

According to another embodiment of the vaccine composition according to the invention, the nucleotide sequence, preferably a DNA, is complexed with the DEAE-dextran (Pagano t al., 1967) or with nuclear proteins (Kaneda et al., 1989), with lipids (Felgner et al., 1987) or encapsulated into liposomes (Fraley et al., 1980) or alternatively introduced in the form of a gel

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facilitating its transfection into the cells (Midoux et al., 1993, Pastore et al., 1994). The polynucleotide or the vector according to the invention may also be in suspension in a buffer solution or may be combined with liposomes.

Advantageously, such a vaccine will be prepared in accordance with the technique described by Tacson et al. or Huygen et al. in 1996 or alternatively in accordance with the technique described by Davis et al. in International Application No. WO 95/11307.

Such a vaccine may also be prepared in the form of a composition containing a vector according to the invention, placed under the control of regulatory elements allowing its expression in humans or animals. It is possible, for example, to use, as vector for the *in vivo* expression of the polypeptide antigen of interest, the plasmid pcDNA3 or the plasmid pcDNA1/neo, both marketed by Invitrogen (R & D Systems, Abingdon, United Kingdom). It is also possible to use the plasmid V1Jns.tPA, described by Shiver et al. in 1995. Such a vaccine will advantageously comprise, in addition to the recombinant vector, a saline solution, for example a sodium chloride solution.

The immunogenic compositions of the invention can be utilized as part of methods of immunization, wherein such methods comprise administering to a host, e.g., a human host, an immunizing amount of the immunogenic compositions of the invention. In a preferred embodiment, the method of immunizing is a method of immunizing against Chlamydia trachomatis.

A pharmaceutically acceptable vehicle is understood to designate a compound or a combination of compounds entering into a pharmaceutical or vaccine composition which does not cause side effects and which makes it possible, for example, to facilitate the administration of the active compound, to increase its life and/or its efficacy in the body, to increase its solubility in solution or alternatively to enhance its preservation. These pharmaceutically acceptable vehicles are well known and will be adapted by persons skilled in the art according to the nature and the mode of administration of the active compound chosen.

As regards the vaccine formulations, these may comprise appropriate immunity adjuvants which are known to persons skilled in the art, such as, for example, aluminium hydroxide, a representative of the family of muramyl peptides such as one of the peptide derivatives of N-acetylmuramyl, a bacterial lysate, or alternatively incomplete Freund's adjuvant, Stimulon<sup>TM</sup> QS-21 (Aquila Biopharmaceuticals, Inc., Framinham, MA), MPL<sup>TM</sup> (3-0-deacylated monophosphoryl lipid A; RIBI ImmunoChem Research, Inc., Hamilton, MT), aluminum phosphate, IL-12 (Genetics Institute, Cambridge, MA).

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Preferably, these compounds will be administered by the systemic route, in particular by the intravenous route, by the intranasal, intramuscular, intradermal or subcutaneous route, or by the oral route. More preferably, the vaccine composition comprising polypeptides according to the invention will be administered several times, spread out over time, by the intradermal or subcutaneous

route.

Their optimum modes of administration, dosages and galenic forms may be determined according to criteria which are generally taken into account in establishing a treatment adapted to a patient, such as for example the patient's age or body weight, the seriousness of his general condition, tolerance of the treatment and the side effects observed.

The invention comprises the use of a composition according to the invention for the treatment or the prevention of genital diseases which are induced or worsened by *Chlamydia trachomatis*.

Finally, the invention comprises the use of a composition according to the invention for the treatment or the prevention of eye diseases which are induced or worsened by the presence of *Chlamydia trachomatis*.

Finally, the invention comprises the use of a composition according to the invention for the treatment or the prevention of systemic diseases, especially of the lymphatic system, which are induced or worsened by the presence of *Chlamydia trachomatis*.

Other characteristics and advantages of the invention appear in the following examples and figures:

## Legend to the figures:

Figure 1: Line for the production of Chlamydia trachomatis sequences

20 Figure 2: Analysis of the sequences and assembling

Figure 3: Finishing techniques

Figure 3a): Assembly map

Figure 3b): Determination and use of the orphan ends of the contigs

#### 25 EXAMPLES

Cells

The Chlamydia trachomatis LGV2 strain used is identified to have over 98% homology with the outer membrane protein sequences omp1 (CHTMOMPA) and omp2 (CHTOMP2A) of the Chlamydia trachomatis serovar L2/434/Bu strain.

The Chlamydia trachomatis LGV2 strain is cultured on mouse fibroblasts (McCoy cells), obtained from the American Type Culture Collection, under the reference ATCC CRL-1696.

## Culture of the cells

The mouse fibroblasts are cultured in 75-ml cell culture flasks (Corning). The culture 35 medium is Dulbecco's modified cell culture medium (Gibco BRL No. 04101965) supplemented with

MEM amino acids (Gibco BRL - No. 04301140) L (5 ml per 500 ml of medium) and 5% foetal calf serum (Gibco BRL No. 10270 batch 40G8260K) without antibiotics or antifungals.

The cell culture stock is maintained in the following manner. The cell cultures are examined under an inverted microscope. 24 hours after confluence, each cellular lawn is washed with 5 PBS (Gibco BRL No. 04114190), rinsed and then placed for 5 min in an oven in the presence of 3 ml of trypsine (Gibco BRL No. 25200056). The cellular lawn is then detached and then resuspended in 120 ml of culture medium, the whole is stirred in order to make the cellular suspension homogeneous. 30 ml of this suspension are then distributed per cell culture flask. The flasks are kept in a CO2 oven (5%) for 48 hours at a temperature of 37°C. The cell stock is maintained so as to have available daily 10 16 flasks of subconfluent cells. It is these subconfluent cells which will be used so as to be infected with Chlamydia. 25-ml cell culture flasks are also used, these flasks are prepared in a similar manner but the volumes used for maintaining the cells are the following: 1 ml of trypsine, 28 ml of culture medium to resuspend the cells, 7 ml of culture medium are used per 25-ml flask.

## Infection of the cells with Chlamydia

Initially, the Chlamydiae are obtained frozen (at -70°C), in suspension in a volume of 1 millilitre. This preparation is slowly thawed, 500 µl are collected and brought into contact with subconfluent cells, which are obtained as indicated above, in a 25-ml cell culture flask, containing 1 ml of medium, so as to cover the cells. The flask is then centrifuged at 2000 rpm in a «swing» rotor for microtitre plates, the centrifuge being maintained at a temperature of 35°C. After centrifugation, the two flasks are placed in an oven at 35°C for three hours. 6 ml of culture medium containing cycloheximide (1 µg/ml) are then added and the flask is stored at 35°C. After 48 hours, the level of infection is evaluated by direct immunofluorescence and by the cytopathogenic effect caused to the cells.

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#### Direct immunofluorescence

Starting with infected cells, which were obtained as indicated above, a cellular smear is deposited with a Pasteur pipette on a microscope slide. The cellular smear is fixed with acetone for 10 minutes; after draining the acetone, the smear is covered with 30 µl of murine monoclonal 30 antibodies directed against MOMP (major outer membrane protein) of Chlamydia (Syva, Biomérieux) labelled with fluorescein isothiocyanate. The whole is then incubated in a humid chamber at a temperature of 37°C. The slides are then rinsed with water, slightly dried, and then after depositing a drop of mounting medium, a coverslip is mounted before reading. The reading is carried out with the aid of a fluorescence microscope equipped with the required filters (excitation at 490 nm, emission at 520 nm).

## Harvesting of the Chlamydia trachomatis

After checking the infection by direct immunofluorescence, carried out as indicated above, the culture flasks are opened under a sterile cabinet, sterile glass beads with a diameter of the order of a millimeter are placed in the flask. The flask is closed and then vigorously stirred while being maintained horizontally, the cellular lawn at the bottom, so that the glass beads can have a mechanical action on the cellular lawn. Most of the cells are thus detached or broken; the effect of the stirring is observed under an optical microscope so as to ensure proper release of Chlamydiae.

#### Large-scale infection of the cell cultures

The product of the Chlamydiae harvest (culture medium and cellular debris) is collected with a pipette, and distributed into three cell culture flasks containing subconfluent L cells, obtained as indicated above. The cells thus inoculated are placed under gentle stirring (swing) in an oven at 35°C. After one hour, the flasks are kept horizontally in an oven so that the culture medium covers the cells for 3 hours. 30 ml of culture medium containing actydione (1 µg/ml) are then added to each of the flasks. The culture flasks are then stored at 35°C for 48 hours. The cells thus infected are examined under an optical microscope after 24 hours, the cytopathogenic effect is evaluated by the appearance of cytoplasmic inclusions which are visible under an inverted optical microscope. After 48 hours, the vacuoles containing the Chlamydiae occupy the cytoplasm of the cell and push the cell nucleus sideways. At this stage, numerous cells are spontaneously destroyed and have left free elementary bodies in the culture medium. The Chlamydiae are harvested as described above and are either frozen at -80°C or used for another propagation.

#### Purification of the Chlamydiae

The product of the Chlamydia harvests, stored at -80°C, is thawed on a water bath at room temperature. After thawing, each tube is vigorously stirred for one minute and immersed for one minute in an ultrasound tank (BRANSON 1200); the tubes are then stirred by inverting before being centrifuged for 5 min at 2000 rpm. The supernatant is carefully removed and kept at cold temperature (ice). The supernatant is vigorously stirred and then filtered on nylon filters having pores of 5 microns in diameter on a support (Nalgene) allowing a delicate vacuum to be established under the nylon filter. For each filtration, three nylon filters are superposed; these filters are replaced after every 40 ml of filtrate. Two hundred milliliters of filtration product are kept at cold temperature, and then after stirring by inverting, are centrifuged at 10,000 rpm for 90 min, the supernatant is removed and the pellet is taken up in 10 ml of 10 mM Tris, vigorously vortexed and then centrifuged at 10,000 rpm for 90 min. The supernatant is removed and the pellet is taken up in a buffer (20 mM Tris pH 8.0, 50 mM

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KCl, 5 mM MgCl<sub>2</sub>) to which 800 units of DNAse I (Boehringer) are added. The whole is kept at 37°C for one hour. One ml of 0.5 M EDTA is then added, and the whole is vortexed and frozen at -20°C.

## Preparation of the DNA

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The Chlamydiae purified above are thawed and subjected to a proteinase K (Boehringer) digestion in a final volume of 10 ml. The digestion conditions are the following: 0.1 mg/ml proteinase K, 0.1 % SDS at 55°C, stirring every 10 min. The product of digestion is then subjected to a double extraction with phenol-chloroform, two volumes of ethanol are added and the DNA is directly recovered with a Pasteur pipette having one end in the form of a hook. The DNA is dried on the edge of the tube and then resuspended in 500  $\mu$ l of 2 mM Tris pH 7.5. The DNA is stored at 4°C for at least 24 hours before being used for the cloning.

#### Cloning of the DNA

After precipitation, the DNA is quantified by measuring the optical density at 260 nm. Thirty  $\mu g$  of Chlamydia DNA are distributed into 10 tubes of 1.5 ml and diluted in 300  $\mu l$  of water. Each of the tubes is subjected to 10 applications of ultrasound lasting for 0.5 sec in a sonicator (Unisonix XL2020). The contents of the 10 tubes are then grouped and concentrated by successive extractions with butanol (Sigma B1888) in the following manner: two volumes of butanol are added to the dilute DNA mixture. After stirring, the whole is centrifuged for five minutes at 2500 rpm and the butanol is removed. This operation is repeated until the volume of the aqueous phase is less than 1 ml. The DNA is then precipitated in the presence of ethanol and of 0.5 M sodium acetate pH 5.4, and then centrifuged for thirty minutes at 15,000 rpm at cold temperature (4°C). The pellet is washed with 75% ethanol, centrifuged for five minutes at 15,000 rpm and dried at room temperature. A tenth of the preparation is analysed on a 0.8% agarose gel. Typically, the size of the DNA fragments thus prepared is between 200 and 8000 base pairs.

To allow the cloning of the DNA obtained, the ends are repaired. The DNA is distributed in an amount of 10 µg/tube, in the following reaction medium: 100 µl final volume, 1 H buffer (Biolabs 201L),  $0.5\,\mu l$  BSA  $0.05\,m g/m l$ ,  $0.1\,m M$  dATP,  $0.1\,m M$  each of dGTP, dCTP or dTTP, 60,000 IU T4 DNA polymerase. The reaction is incubated for thirty minutes at 16°C. The 30 contents of each of the tubes are then grouped before carrying out an extraction with phenolchloroform and then precipitating the aqueous phase as described above. After this step, the DNA thus prepared is phosphorylated. For that, the DNA is distributed into tubes in an amount of 10 µg per tube, and then in a final volume of 50 µl, the reaction is prepared in the following manner: 1 mM ATP, 1 x kinase buffer, 10 IU T4 polynucleotide kinase (Biolabs 201L). The preparation is incubated for thirty minutes at 37°C. The contents of the tubes are combined and a phenol-chloroform extraction

and then a precipitation are carried out in order to precipitate the DNA. The latter is then suspended in 1 µl of water and then the DNA fragments are separated according to their size on a 0.8% agarose gel (1 x TAE). The DNA is subjected to an electric field of 5 V/cm and then visualized on a UV table. The fragments whose size varies between 1200 and 2000 base pairs are selected by cutting out the gel.

5 The gel fragment thus isolated is placed in a tube and then the DNA is purified with the Qiaex kit (20021 Qiagen), according to the procedure provided by the manufacturer.

#### Preparation of the vector

14 μg of the cloning vector pGEM-5Zf (Proméga P2241) are diluted in a final volume of 150 μl and are subjected to digestion with the restriction enzyme EcoRV 300 IU (Biolabs 195S) according to the protocol and with the reagents provided by the manufacturer. The whole is placed at 37°C for 150 min and then distributed in the wells of a 0.8% agarose gel subjected to an electric field of 5 V/cm. The linearized vector is visualized on a UV table, isolated by cutting out the gel and then purified by the Qiaex kit (Qiagen 20021) according to the manufacturer's recommendations. The purification products are grouped in a tube, the volume is measured and then half the volume of phenol is added and the whole is vigorously stirred for 1 min. Half the volume of chloroform-isoamyl alcohol 24:1 is added and vigorously stirred for 1 min. The whole is centrifuged at 15,000 rpm for 5 min at 4°C, the aqueous phase is recovered and transferred into a tube. The DNA is precipitated in the presence of 0.3 M sodium acetate, pH 5.4 and 3 volumes of ethanol and placed at -20°C for 1 hour. The DNA is then centrifuged at 15,000 rpm for 30 min at 4°C, the supernatant is removed while preserving the pellet, washed twice with 70% ethanol. After drying at room temperature, the DNA is suspended in 25 μl of water.

#### Phosphorylation of the vector

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 $25~\mu l$  of the vector prepared in the preceding step are diluted in a final volume of 500  $\mu l$  of the following reaction mixture:

After repair, the DNA is subjected to a phenol-chloroform extraction and a precipitation, the pellet is then taken up in 10 µl of water, the DNA is quantified by measuring the optical density at 260 nm. The quantified DNA is ligated into the vector PGEm-5Zf(+) prepared by the restriction enzyme EcoRV and dephosphorylated (see preparation of the vector). The ligation is carried out under three conditions which vary in the ratio between the number of vector molecules and the number of insert molecules. Typically, an equimolar ratio, a ratio of 1:3 and a ratio of 3:1 are used for the ligations which are, moreover, carried out under the following conditions: vector PGEm-5Zf(+) 25 ng, cut DNA, ligation buffer in a final volume of 20 µl with T4 DNA ligase (Amersham E70042X); the whole is then placed in a refrigerator overnight and then a phenol-

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chloroform extraction and a precipitation are carried out in a conventional manner. The pellet is taken up in 5 µl of water.

#### Transformation of the bacteria

Plating of the bacteria

Petri dishes containing LB Agar medium containing ampicillin (50  $\mu$ g/ml), Xgal (280  $\mu$ g/ml) [5-bromo-4-chloro-indolyl-beta-D-galactopyranoside (Sigma B-4252)], IPTG (140  $\mu$ g/ml) [isopropyl-beta-D-thiogalactoside (Sigma I-6758)] are used, 50 and 100  $\mu$ l of bacteria are plated for each of the ligations. The Petri dishes are placed upside down at 37°C for 15 to 16 hours in an oven. The number of «recombinant» positive clones is evaluated by counting the white colonies and the blue colonies which are thought to contain the vector alone.

#### Evaluation of the «recombinant» positive clones:

Ninety-four white colonies and two blue colonies are collected with the aid of sterile cones and are deposited at the bottom of the wells of plates designed for carrying out the amplification techniques. 30 µl of the following reaction mixture are added to each well: 1.7 mM MgCl<sub>2</sub>, 0.2 mM each of dATP, dCTP, dGTP and dTTP, two synthetic oligonucleotides corresponding to sequences flanking the cloning site on either side and orienting the synthesis of the DNA in a convergent manner (0.5 µM RP and PU primers, 1 U TAQ polymerase (GibcoBRL 18038-026)).

The colonies thus prepared are subjected to a temperature of 94°C for 5 min and then to 30 thermal cycles composed of the following steps: 94°C for 40 s, 50°C for 30 s, 72°C for 180 s. The reaction is then kept for 7 min at 72°C and then kept at 4°C.

The amplification products are deposited on an agarose gel (0.8%), stained with ethidium bromide, subjected to electrophoresis, and then analysed on an ultraviolet table. The presence of an amplification fragment having a size greater than 500 base pairs indicates the presence of an insert. The bacterial clones are then prepared so as to study the sequence of their insert.

#### Sequencing

To sequence the inserts of the clones obtained as above, these were amplified by PCR on bacteria cultures carried out overnight using the primers for the vectors flanking the inserts. The sequence of the ends of these inserts (on average 500 bases on each side) was determined by automated fluorescent sequencing on an ABI 377 sequencer, equipped with the ABI Prism DNA Sequencing Analysis software (version 2.1.2).

The sequences obtained by sequencing in a high-yield line (Figure 1) are stored in a database; this part of the production is independent of any treatment of the sequences. The sequences are extracted from the database, avoiding all the regions of inadequate quality, that is to say the regions for which uncertainties are observed on the sequence at more than 95%. After extraction, the sequences are introduced into a processing line, the diagram of which is described in Figure 2. In a first path of this processing line, the sequences are assembled by the Gap4 software from R. Staden (Bonfield et al., 1995) (OS UNIX/SUN Solaris); the results obtained by this software are kept in the form of two files which will be used for a subsequent processing. The first of these files provides information on the sequence of each of the contigs obtained. The second file represents all the clones participating in the composition of all the contigs as well as their positions on the respective contigs.

The second processing path uses a sequence assembler (TIGR-Asmg assembler UNIX/SUN Solaris); the results of this second processing path are kept in the form of a file in the TIGR-Asmg format which provides information on the relationship existing between the sequences selected for the assembly. This assembler is sometimes incapable of linking contigs whose ends 15 overlap over several hundreds of base pairs.

The results obtained from these two assemblers are compared with the aid of the BLAST program, each of the contigs derived from one assembly path being compared with the contigs derived from the other path.

For the two processing paths, the strict assembly parameters are fixed (95% 20 homology, 30 superposition nucleotides). These parameters avoid 3 to 5% of the clones derived from eukaryotic cells being confused with sequences obtained from the clones derived from Chlamydia trachomatis. The eukaryotic sequences are however preserved during the course of this project; the strategy introduced, which is described below, will be designed, inter alia, not to be impeded by these sequences derived from contaminating clones.

The results of these two assemblers are processed in a software developed for this project. This software operates on a Windows NT platform and receives, as data, the results derived from the STADEN software and/or the results derived from the TIGR-Asmg assembler, the software, results, after processing of the data, in the determination of an assembly map which gives the proximity relationship and the orientation of the contigs in relation to one another (Figure 3a). Using 30 this assembly map, the software determines all the primers necessary for finishing the project. This treatment, which will be detailed below, has the advantage of distinguishing the isolated sequences derived from the contaminations, by the DNA eukaryotic cells, of the small-sized sequences clearly integrated into the project by the relationships which they establish with contigs. In order to allow, without any risk of error, the arrangement and the orientation of the contigs in relation to one another, a statistical evaluation of the accuracy of the names «naming» of sequence is made from the results of

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«contigation». This evaluation makes it possible to give each of the clone plates, as well as each of the subsets of plates, a weight which is inversely proportional to probable error rate existing in the «naming» of the sequences obtained from this plate or from a subset of this plate. In spite of a low error rate, errors may occur throughout the steps of production of the clones and of the sequences. These steps are numerous, repetitive and although most of them are automated, others, like the deposition in the sequencers, are manual; it is then possible for the operator to make mistakes such as the inversion of two sequences. This type of error has a repercussion on the subsequent processing of the data, by resulting in relationships (between the contigs) which do not exist in reality, then in attempts at directed sequencing between the contigs which will end in failure. It is because of this that 10 the evaluation of the naming errors is of particular importance since it allows the establishment of a probabilistic assembly map from which it becomes possible to determine all the clones which will serve as template to obtain sequences separating two adjacent contigs. Table 2 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997, each of which is 15 incorporated by reference herein in its entirety, gives the clones and the sequences of the primers initially used during the initial operations.

To avoid the step which consists in ordering and then preparing the clones by conventional microbiological means, outer and inner primers oriented towards the regions not yet sequenced are defined by the software. The primers thus determined make it possible to prepare, by 20 PCR, a template covering the nonsequenced region. It is the so-called outer primers (the ones most distant from the region to be sequenced) which are used to prepare this template. The template is then purified and a sequence is obtained on each of the two strands during 2 sequencing reactions which each use one of the 2 inner primers. In order to facilitate the use of this approach, the two outer primers and the two inner primers are prepared and then stored on the same position of 4 different 25 96-well plates. The two plates containing the outer primers are used to perform the PCRs which serve to prepare the templates. These templates will be purified on purification columns preserving the topography of the plates. Each of the sequences are obtained using primers situated on one and then on the other of the plates containing the inner primers. This distribution allows a very extensive automation of the process and results in a method which is simple to use for finishing the regions not yet sequenced. Table 3 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997, each of which is incorporated by reference herein in its entirety, gives the names and the sequences of the primers used for finishing Chlamydia trachomatis.

Finally, a number of contigs exist in a configuration where one of their ends is not linked to any other contig end (Figure 3b) by a connecting clone relationship (a connecting clone is

defined as a clone having one sequence end on a contig and the other end of its sequence on another contig; furthermore, this clone must be derived from a plate or a subset of plates with adequate naming quality). For the Chlamydia trachomatis project, this particular case occurred 37 times. Two adjacent PCR primers orienting the synthesis of the DNA towards the end of the consensus sequence are defined for each of the orphan ends of the consensus sequence. The primer which is closest to the end of the sequence is called the inner primer whereas the primer which is more distant from the end of the sequence is called the outer primer. The outer primers are used to explore the mutual relationship between the orphan ends of the different contigs. The presence of a single PCR product and the possibility of amplifying this product unambiguously using the inner primers evokes the probable relationship between the contigs on which the primers which allowed the amplification are situated. This relationship will be confirmed by sequencing and will allow the connection between the orphan ends of the consensus sequences. This strategy has made it possible to obtain a complete map of the Chlamydia trachomatis chromosome and then to finish the project.

### 15 Quality control

All the bases not determined with certainty in the chromosomal sequence were noted and the density of uncertainties was measured on the entire chromosome. The regions with a high density of uncertainties were noted and the PCR primers spanning these regions were drawn and are represented in Table 4 of parent U.S. Application Serial No. 60/107077 filed November 4, 1998, French application 97-15041 filed November 28, 1997 and French application 97-16034 filed December 17, 1997 each of which is incorporated by reference herein in its entirety.

#### Data banks

Local reorganizations of major public banks were used. The protein bank used consists of the nonredundant fusion of the Genpept bank (automated translation of GenBank, NCBI; Benson et al., 1996).

The entire BLAST software (public domain, Altschul et al., 1990) for searching for homologies between a sequence and protein or nucleic data banks was used. The significance levels used depend on the length and the complexity of the region tested as well as the size of the reference bank. They were adjusted and adapted to each analysis.

The results of the search for homologies between a sequence according to the invention and protein or nucleic data banks are presented and summarized in Table 1 below.

Table 1: <u>List of coding chromosome regions and homologies between these regions and the sequence</u>

55 banks.

Legend to Table 1: Open reading frames are identified with the GenMark software version 2.3A (GenePro), the template used is *Chlamydia trachomatis* of order 4 on a length of 196 nucleotides with a window of 12 nucleotides and a minimum signal of 0.5. These reading frames are numbered in order of appearance on the chromosome, starting with ORF2 (ORF column). The positions of the beginning and of the end are then given in column 2 (position). When the position of the beginning is greater than the position of the end, this means that the region is encoded by the strand complementary to the sequence which was given in the sequence SEQ ID No. 1.

All the putative products were subjected to a search for homology on GENPEPT (release 103 for SEQ ID No. 2 to SEQ ID No. 1076 and release 108 for SEQ ID No. 1077 to SEQ ID No. 1197 with the BLASTp software (Altschul et al. 1990), with, as parameters, the default parameters with the exception of the expected value E set at 10<sup>-5</sup> (for SEQ ID No. 2 to SEQ ID No. 1076) and P value set at e<sup>-10</sup> (for SEQ ID No. 1077 to SEQ ID No. 1197). Subsequently, only the identities greater than 30% (I% column) were taken into account. The description of the most homologous sequence is given in the Homology column; the identifier for the latter sequence is given in the ID column and the animal species to which this sequence belongs is given in the Species column. The Homology score is evaluated by the sum of the blast scores for each region of homology and reported in the Score column. Table 1 also reflects data from additional ORF finder programs as defined below.

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## Materials and methods: transmembrane domains:

The DAS software was used as recommended by the authors (Cserzo et al., 1997).

This method uses, to predict the transmembrane domains, templates derived from a sampling of selected proteins. All the regions for which a "Cutoff" greater than 1.5 was found by the program were taken into account.

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#### Additional ORF Finder Programs

For this analysis, two additional ORF finder programs were used to predict potential open reading frames of a minimum length of 74 amino acids; Glimmer (Salzberg, S.L., Delcher, A., Kasif, S., and W. White. 1998. Microbial gene identification using interpolated Markov models.

Nucleic Acids Res. 26:544-548.), and an in-house written program. The in-house program used a very simple search algorithm. The analysis required the that the genomic DNA sequence text be in the 5' to 3' direction, the genome is circular, and that TAA, TAG, and TGA are stop codons. The search parameters were as follows:

(1) A search for an ORF that started with a GTG codon was performed. If no GTG codons were found, then a search for an ATG codon was performed. However, if a GTG codon was found,

then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.

- (2) A search for an ORF that started with a TTG codon was performed. If no TTG codons were found, then a search for a ATG codon was performed. However, if a TTG codon was found, 5 then a search downstream for a ATG codon was performed. All start and stop nucleotide positions were recorded.
  - (3) The analysis described in steps 1 and 2 were repeated for the opposite strand of DNA sequence.
- (4) A search for ORFs that determined all ORF lengths using start and stop positions in the same reading frames was performed.
  - (5) All ORFs whose DNA length was less than 225 nucleotides were eliminated from the search.

## Surface Exposed Protein Search Criteria

Potential cell surface vaccine targets are outer membrane proteins such as porins, lipoproteins, adhesions and other non-integral proteins. In Chlamydia psittaci, the major immunogens is a group of putative outer membrane proteins (POMPs) and no homologs have been found in Chlamydia trachomatis and Chlamydia trachomatis by traditional analysis (Longbottom, D., Russell, M., Dunbar, S.M., Jones, G.E., and A.J. Herring. 1998. Molecular Cloning and 20 Characterization of the Genes Coding for the Highly Immunogenic Cluster of 90-Kilodalton Envelope Proteins from Chlamydia psittaci Subtype That Causes Abortion in Sheep. Infect Immun 66:1317-1324.) However, utilizing the criteria described below, several ORFs encoding outer membrane proteins have been identified in Chlamydia trachomatis, all of which may represent vaccine candidates. Any ORF which met any one of the criteria described below were considered to encode a 25 surface exposed protein.

Protein homology searches of the translated ORFs were done using the Blastp 2.0 tool (Altschul, S.F., Madden, T.L., Schaffer, A.A., Zhang, J., Zhang, Z., Miller, W., and D.J. Lipman. 1997. Gapped BLAST and PSI-BLAST: a new generation of protein database search programs. Nucleic Acids Res. 25:3389-3402). An ORF was labeled surface exposed if the translated ORF had 30 homology to a known, or hypothetical, or putative surface exposed protein with a P score less than e<sup>-10</sup>.

Most, if not all, proteins that are localized to the membrane of bacteria, via a secretory pathway, contain a signal peptide. The software program SignalP, was used to analyze the amino acid sequence of an ORF for such a signal peptide (Nielsen, H., Engelbrecht. J., Brunak, S., 35 and G. von Heijne. 1997. Identification of prokaryotic and eukaryotic signal peptides and prediction

of their cleavage sites. Protein Engineering 10:1-6.) The first 60 N-terminal amino acids of each ORF were analyzed by SignalP using the Gram-negative software database. The output generates four separate values, maximum C, maximum Y, maximum S, and mean S. The S-score, or signal region, is the probability of the position belonging to the signal peptide. The C-score, or cleavage site, is the probability of the position being the first in the mature protein. The Y-score is the geometric average of the C-score and a smoothed derivative of the S-score. A conclusion of either a Yes or No is given next to each score. If all four conclusions are Yes and the C-terminal amino acid is either a phenylalanine (F) or a tyrosine (Y), the ORF was labelled outer membrane (Struyve, M., Moons, M., and J. Tommassen. 1991. Carboxy-terminal Phenylalanine is Essential for the Correct Assembly of a Bacterial Outer Membrane Protein. J. Mol. Biol. 218:141-148.)

The program called Psort was used to determine the localization of a protein based on its signal sequence, recognition of transmembrane segments, and analysis of its amino acid composition (Nakai, K., and M. Kanehisa. 1991. Expert system for predicting protein localization sites in gram-negative bacteria. Proteins 11:95-110.) An ORF is considered to be an outer membrane protein if the output data predicts the ORF encoded protein as outer membrane with a certainty value of 0.5 or better and whose value is at least twice as large as the next predicted localized certainty value.

Finally, ORFs that were not predicted to be outer membrane or surface exposed, based on the above criteria, were further analyzed. The Blastp output data for these ORFs were searched using various general and specific keywords, suggestive of known cell surface exposed proteins. An ORF was labeled surface exposed if the keywords matched had a Blastp hit with a P score less than e<sup>-10</sup>, and there was no better data indicating otherwise. The following is a list of the searched keywords:

25	Adhesion	Adhesin	Invasin
	Invasion	Extension	Omp
	Outer Surface	Porin	Outer Membrane
	Cell Surface	Cell Wall	Pilin
	Flagellar sheath	Cir	ChuA
30	СорВ	ExeD	FadL
	FecA	FepA	FhuA
	FmdC	FomA	FrpB
	GspD	HemR	HgbA
	Hgp	HmbR	HmuR
35	HMW	HrcC	Hrp

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	InvG	LamB	LbpA
	LcrQ	Lmp1	MxiD
	MOMP	PilE	HpaA
	NolW	NspA	OpcP
5	OpnP	Opr	OspA
	PhoE	PldA	Por
	PscC	PulD	PupA
	QuiX	RafY	ScrY
	SepC	ShuA	SomA
10	SpiA	Tbpl	Yop
	YscC	mip	Tol
	Pilus	BtuB	

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Those ORFs that did not meet the minimum requirement for being an outer membrane protein based on the above search criteria but which were homologous to identified outer membrane ORFs in Chlamydia pneumoniae were included. The Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) was analyzed using the above search criteria and a number of outer membrane ORFs were identified. These Chlamydia pneumoniae ORFs were then tested against the Chlamydia trachomatis genome using Blastp. Any Chlamydia trachomatis ORF with a Blastp P value less than e<sup>-10</sup> against a Chlamydia pneumoniae outer membrane was included in this section, if there was no better data indicating otherwise. A list of ORFs in the Chlamydia trachomatis genome encoding putative surface exposed proteins is set forth above in the specification.

# Identification of Putative Lipoproteins in the Genome of Chlamydia trachomatis

Lipoproteins are the most abundant post-translationally modified bacterial secretory proteins (Pugsley, A. P., 1993). The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108). The characteristic features of lipoproteins are a thiol-linked diacylglyceride and an amine-linked monoacyl group on the cysteine that becomes the amino-terminal residue after signal peptide cleavage by Signal Peptidase II. (Pugsley, A. P., 1993). The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108). The identification of putative lipoproteins from the genomic sequencing of Chlamydia trachomatis was done by examining the deduced amino acid sequence of identified ORFs for the presence of a signal peptide with a Signal Peptidase II cleavage site analogous to the consensus sequence for prolipoprotein modification and processing reactions (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York;

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Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128).

The deduced amino acid sequences of *Chlamydia trachomatis* ORFs were initially screened for the most basic of lipoprotein characteristics, a cysteine in the first 30 amino acids of the deduced protein. ORFs with a standard start codon (ATG, GTG, or TTG) and having one or more of the following characteristics were selected for direct analysis of their first 30 amino acids:

- (a) Significant Signal P value (at least two out-of-the four values are Yes)
- (b) PSORT value indicating membrane passage (IM-inner membrane, Peri-periplasm, or OM-outer membrane)
  - (c) Identification of the word lipoprotein among the ORF Blastp data set.
- (d) A Blastp value of <e<sup>-10</sup> with a putative lipoprotein from *Chlamydia pneumoniae* (French application No. 97-14673 filed 21 November 1997).

The first 30 amino acids encoded by each ORF in this set were analyzed for the characteristics commonly found in lipoprotein signal peptides (Pugsley, A. P., 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108; Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York; Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Grampositive bacteria. J. Bacteriol. 177:1123-1128.) Putative lipoprotein signal peptides were required to have a cysteine between amino acid 10 and 30 and reach a minimum score of three based on the following criteria for lipoprotein signal peptides:

- (a) Identification of specific amino acids in specific positions around the cysteine which are part of the consensus Signal Peptidase II cleavage site (Hayashi, S., and H. C. Wu. 1992. Identification and characterization of lipid-modified proteins in bacteria, p. 261-285. In N. M. Hooper and A. J. Turner (ed.) Lipid modification of proteins: A practical approach. Oxford University Press, New York); Sutcliffe, I. C. and R. R. B. Russell. 1995. Lipoproteins of Gram-positive bacteria. J. Bacteriol. 177:1123-1128). Since the identification of the cleavage site is the most important factor in identifying putative lipoproteins, each correctly positioned amino acid contributed toward reaching the minimum score of three.
  - (b) A hydrophobic region rich in alanine and leucine prior to the cleavage site (Pugsley, A. P., 1993. The complete general secretory pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.
- (c) A short stretch of hydrophilic amino acids greater than or equal to 1, usually lysine or arginine, following the N-terminal methionine (Pugsley, A. P., 1993. The complete general secretory

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pathway in Gram-negative bacteria. Microbiol. Rev. 57:50-108) contributed toward reaching the minimum score of three.

A list of ORFs in the *Chlamydia trachomatis* genome encoding putative lipoproteins is set forth above in the specification.

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## LPS-Related ORFs of Chlamydia trachomatis

Lipopolysaccharide (LPS) is an important major surface antigen of Chlamydia cells. Monoclonal antibodies (Mab) directed against LPS of Chlamydia pneumoniae have been identified that can neutralize the infectivity of Chlamydia pneumoniae both in vitro and in vivo (Peterson et al. 10 1988). Similar results are expected utilizing monoclonal antibodies against LPS of Chlamydia trachomatis. LPS is composed of lipid A and a core oligosaccharide portion and is phenotypically of the rough type (R-LPS) (Lukacova, M., Baumann, M., Brade, L., Mamat, U., Brade, H. 1994. Lipopolysaccharide Smooth-Rough Phase Variation in Bacteria of the Genus Chlamydia. Infect. Immun. June 62(6):2270-2276.) The lipid A component is composed of fatty acids which serve to anchor LPS in the outer membrane. The core component contains sugars and sugar derivatives such as a trisaccharide of 3-deoxy-D-manno-octulosonic acid (KDO) (Reeves, P.R., Hobbs, M., Valvano, M.A., Skurnik, M., Whitfield, C., Coplin, D., Kido, N., Klena, J., Maskell, D., Raetz, C.R.H., Rick, P.D. 1996. Bacterial Polysaccharide Synthesis and Gene Nomenclature pp. 10071-10078, Elsevier Science Ltd.). The KDO gene product is a multifunctional glycosyltransferase and represents a 20 shared epitope among the Chlamydia. For a review of LPS biosynthesis see, e.g., Schnaitman, C.A., Klena, J.D. 1993. Genetics of Lipopolysaccharide Biosynthesis in Enteric Bacteria. Microbiol. Rev. 57:655-682.

A text search of the ORF Blastp results identified several genes that are involved in Chlamydial LPS production with a P score less than e<sup>-10</sup>. The following key-terms were used in the text search: KDO, CPS (Capsular Polysaccharide Biosynthesis), capsule, LPS, rfa, rfb, rfc, rfe, rha, rhl, core, epimerase, isomerase, transferase, pyrophosphorylase, phosphatase, aldolase, heptose, manno, glucose, lpxB, fibronectin, fibrinogen, fucosyltransferase, lic, lgt, pgm, tolC, rol, ChoP, phosphorylcholine, waaF, PGL-Tb1. A list of ORFs in the Chlamydia trachomatis genome encoding putative polypeptides involved in LPS biosynthesis is set forth above in the specification.

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#### Type III And Other Secreted Products

Type III secretion enables gram-negative bacteria to secrete and inject pathogenicity proteins into the cytosol of eukaryotic host cells (Hueck, C. J., 1998. Type III Protein Secretion Systems in Bacterial Pathogens of Animals and Plants. In Microbiology and Molecular Biology Reviews. 62:379-433.) These secreted factors often resemble eukaryotic signal transduction factors,

thus enabling the bacterium to redirect host cell functions (Lee, C.A., 1997. Type III secretion systems: machines to deliver bacterial proteins into eukaryotic cells? Trends Microbiol. 5:148-156.) In an attempt to corrupt normal cellular functions, Chlamydial pathogenicity factors injected into the host cytosol will nonetheless, as cytoplasmic constituents be processed and presented in the context of the Major Histocompatibility Complex (MHC class I). As such, these pathogenicity proteins represent MHC class I antigens and will play an important role in cellular immunity. Also included in this set are secreted non-type III products that may play a role as vaccine components.

A text search of the ORF Blastp results identified genes that are involved in Chlamydia trachomatis protein secretion with a P score less than e-10. The following key-terms were 10 used in the text search in an effort to identify surface localized or secreted products: Yop, Lcr, Ypk, Exo, Pcr, Pop, Ipa, Vir, Ssp, Spt, Esp, Tir, Hrp, Mxi, hemolysin, toxin, IgA protease, cytolysin, tox, hap, secreted and Mip.

Chlamydia trachomatis ORFs that did not meet the above keyword search criteria, but have homologs in Chlamydia pneumoniae that do meet the search criteria are included herein. The Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) was analyzed using the above search criteria and a number of ORFs were identified. These Chlamydia pneumoniae ORFs were tested against the Chlamydia trachomatis genome using Blastp. Any Chlamydia trachomatis ORF with a Blastp P value < e.10 against a Chlamydia pneumoniae homolog, identified using the above search criteria, was included. A list of ORFs in the Chlamydia 20 trachomatis genome encoding putative secreted proteins is set forth above in the specification.

# Chlamydia trachomatis RGD Recognition Sequence

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Proteins that contain Arg-Gly-Asp (RGD) attachment site, together with integrins that serve as their receptor constitute a major recognition system for cell adhesion. The RGD sequence is the cell attachment site of a large number of adhesive extracellular matrix, blood, and cell surface proteins and nearly half of the known integrins recognize this sequence in their adhesion protein ligands. There are many RGD containing microbial proteins such as the penton protein of adenovirus, the coxsackie virus, the foot and mouth virus and pertactin, a 69 kDa (kilodalton) surface protein of Bordetella pertussis, that serve as ligands through which these microbes bind to integrins on the cell 30 surfaces and gain entry into the cell. The following provides evidence supporting the importance of RGD in microbial adhesion:

a) The adenovirus penton base protein has a cell rounding activity and when penton base was expressed in E. coli, it caused cell rounding and cells adhered to polystyrene wells coated with the protein. Mutant analysis showed that both these properties required an RGD sequence. Virus mutants with amino acid substitutions in the RGD sequence, showed much less adherence to HeLa S3 cells, and also were delayed in virus reproduction (Bai, M., Harfe, B., and Freimuth, P. 1993. Mutations That Alter an RGD Sequence in the Adenovirus Type 2 Penton Base Protein Abolish Its Cell-Rounding Activity and Delay Virus Reproduction in Flat Cells. J. Virol. 67:5198-5205).

b) It has been shown that attachment and entry of coxsackie virus A9 to GMK cells were dependent on an RGD motif in the capsid protein VP1. VP1 has also been shown to bind  $\alpha_{\nu}\beta_{3}$  integrin, which is a vitronectin receptor (Roivainen, M., Piirainen, L., Hovi, T., Virtanen, I., Riikonen, T., Heino, J., and Hyypia, T. 1994. Entry of Coxsackievirus A9 into Host Cells: Specific Interactions with  $\alpha_{\nu}\beta_{3}$  Integrin, the Vitronectin Receptor Virology, 203:357-65).

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- c) During the course of whooping cough, *Bordetella pertussis* interacts with alveolar macrophages and other leukocytes on the respiratory epithelium. Whole bacteria adheres by means of two proteins, filamentous hemagglutinin (FHA) and pertussis toxin. FHA interacts with two classes of molecules on macrophages, galactose containing glycoconjugates and the integrin CR3. The interaction between CR3 and FHA involves recognition of RGD sequence at the positions 1097-1099 in FHA (Relman, D., Tuomanen, E., Falkow, S., Golenbock, D. T., Saukkonen, K., and Wright, S. D. "Recognitition of a Bacterial Adhesin by an Integrin: Macrophage CR3 Binds Filamentous Hemagglutinin of Bordetella Pertussis." Cell, 61:1375-1382 (1990)).
- d) Pertactin, a 69 kDa outer membrane protein of *Bordetella pertussis*, has been shown to promote attachment of Chinese hamster ovary cells (CHO). This attachment is mediated by recognition of RGD sequence in pertactin by integrins on CHO cells and can be inhibited by synthetic RGD containing peptide homologous to the one present in pertactin (Leininger, E., Roberts, M., Kenimer, J. G., Charles, I. G., Fairweather, N., Novotny, P., and Brennan, M. J. 1991. Pertactin, an Arg-Gly-Asp containing *Bordetella pertussis* surface protein that promotes adherence of mammalian cells Proc. Natl. Acad. Sci. USA, 88:345-349).
- e) The RGD sequence is highly conserved in the VP1 protein of foot and mouth disease virus (FMDV). Attachment of FMDV to baby hamster kidney cells (BHK) has been shown to be mediated 30 by VP1 protein via the RGD sequence. Antibodies against the RGD sequence of VP1 blocked attachment of virus to BHK cells (Fox, G., Parry, N. R., Barnett, P. V., McGinn, B., Rowland, D. J., and Brown, F. 1989. The Cell Attachment Site on Foot-and-Mouth Disease Virus Includes the Amino Acid Sequence RGD (Arginine-Glycine-Aspartic Acid) J. Gen. Virol., 70:625-637).

It has been demonstrated that bacterial adherence can be based on interaction of a bacterial adhesin RGD sequence with an integrin and that bacterial adhesins can have multiple

binding site characteristic of eukaryotic extracellular matrix proteins. RGD recognition is one of the important mechanisms used by microbes to gain entry into eukaryotic cells.

The complete deduced protein sequence of the Chlamydia trachomatis genome was searched for the presence of RGD sequence. There were a total of 38 ORFs that had one or more RGD sequences. Not all RGD containing proteins mediate cell attachment. It has been shown that RGD containing peptides that have proline immediately following the RGD sequence are inactive in cell attachment assays (Pierschbacher & Ruoslahti. 1987. Influence of stereochemistry of the sequence Arg-Gly-Asp-Xaa on binding specificity in cell adhesion. J. Biol. Chem. 262:17294-98). ORFs that had RGD, with proline as the amino acid following the RGD sequence were excluded from the list. Also, RGD sequence may not be available at the surface of the protein or may be present in a context that is not compatible with integrin binding. Since not all RGD-containing proteins are involved in cell attachment, several other criteria were used to refine the list of RGD-containing proteins. A list of ORFs in the Chlamydia trachomatis genome encoding polypeptides with RGD recognition sequence(s) is set forth above in the specification.

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#### Non-Chlamydia pneumoniae ORFs

Chlamydia trachomatis ORFs were compared to the ORFs in the Chlamydia pneumoniae genome (French patent application No. 97-14673, filed 21 November 1997) using Blastp. Any Chlamydia trachomatis ORF with a Blastp «P» value greater than e<sup>-10</sup> (i.e. >e<sup>-10</sup>) against 20 Chlamydia pneumoniae ORFs are included in this section. A list of ORFs in the Chlamydia trachomatis genome which are not found in Chlamydia pneumoniae is set forth above in the specification.

# Cell Wall Anchor Surface ORFs

Many surface proteins are anchored to the cell wall of Gram-positive bacteria via the conserved LPXTG motif (Schneewind, O., Fowler, A., and Faull, K.F. 1995. Structure of the Cell Wall Anchor of Surface Proteins in Staphylococcus aureus. Science 268:103-106). A search of the proteins encoded by the Chlamydia trachomatis ORFs was done using the motif LPXTG. A list of ORFs in the Chlamydia trachomatis genome encoding polypeptides anchored to the cell wall is set forth above in the specification.

#### **ECACC Deposits**

Samples of Chlamydia trachomatis were deposited with the European Collection of Cell Cultures (ECACC), Salisbury, Wiltshire SP4 OJG, UK on November 26, 1998 and assigned the provisional accession number 98112618. Cells can be grown, harvested and purified, and DNA can

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be prepared as discussed above. In order to enable recovery of specific fragments of the chromosome, one can run targeted PCR reactions, whose amplification products can then be sequenced and/or cloned into any suitable vector, according to standard procedures known to those skilled in the art.

In addition, a pool of clones covering the *Chlamydia trachomatis* genome was deposited with the ECACC on November 26, 1998 and assigned provisional accession number 98112617. The pool of clones contains a series of clones, which when taken together, cover the whole chromosome, with a redundancy of slightly more than ten. The total number of clones in the sample is 13,572.

Table 4 lists groups of oligonucleotides to be used to amplify each of ORFs 2-1197 according to standard procedures known to those skilled in the art. Such oligonucleotides are listed as SEQ ID Nos. 1198 to 5981. For each ORF, the following is listed: one forward primer positioned 2,000 bp upstream of the beginning of the ORF; one forward primer positioned 200 bp upstream of the beginning of the ORF; one reverse primer positioned 2,000 bp downstream at the end of ORF, which is 2,000 bp upstream of the end site of the ORF on the complementary strand; and one reverse primer 200 bp downstream at the end of ORF, which is 200 bp upstream of the end site of the ORF on the complementary strand. The corresponding SEQ ID Nos. for the primers are listed in Table 4, where Fp is the proximal forward primer; Fd is the distal forward primer; Bp is the proximal reverse primer; and Bd is the distal reverse primer. The positions of the 5' ends of each of these primers on the nucleotide sequence of SEQ ID No. 1 are shown in Table 5.

The present invention is not to be limited in scope by the specific embodiments described herein, which are intended as single illustrations of individual aspects of the invention, and functionally equivalent methods and components are within the scope of the invention. Indeed, various modifications of the invention, in addition to those shown and described herein will become apparent to those skilled in the art from the foregoing description and accompanying drawings. Such modifications are intended to fall within the scope of the appended claims.

All publications and patent applications mentioned in this specification are herein incorporated by reference to the same extent as if each individual publication or patent application was specifically and individually indicated to be incorporated by reference.

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	Score	2000	379	285	552	265	174	153	210	324	152	000	367	410	324	577	223	573	86	, 643	2214
	Species		Chlamydia psittaci	Haemophilus influenzae	Escherichia coli	1	Staphylococcus carnosus	Bacillus subtilis	Synechococciis sp	1	Methanococcus jannaschij	Sympohocystis		Oryctolagus cuniculus	Saccharopolyspora erythraea	Bacillus subtilis	Escherichia coli		Borrelia burgdorferi	Shidella flexmeri	l est
	£		U72499	U32786	AE000123	U32770	X78084	AF008220	D28752	280108	U67605	D90913	U32691	U75284	L38646	M97391	U18997		U80956	D11024	X52175
TABLE 1	Homology	putative	putative 98 kDa outer membrane protein	lipid A disaccharide synthetase (lpxB)	poly(A) polymerase	D-alanine permease (dagA)	signalpeptidase II	YteA	ORF 168	unknown	hypothetical protein (SP:P39587)	rRNA methylase	hypothetical	neutral amino acid transporter B0.	dihydrolipoamide acetyltransferase	branched chain alpha-keto acid dehydrogenase E2	ORF_0328	putative	putative outer membrane protein	ORF-2	dnaK like protein (AA 1-660)
	stop	208	505	3242	5126	6619	8082	8591	8979	10430	11254	11916	13324	14413	61051	15969	16501	16138	17417	18437	20814
	begin	501	3276	5068	6400	7977	8582	8995	9440	9828	10367	11245	12068	13532	14807	14932	15995	16467	18190	20521	22202
	ORF	ORF2	ORF3	ORF4	ORF5	ORF6	ORF7	ORF8	ORF9	ORF10	ORF11	ORF12	ORF13	ORF14	ORF15	ORF16	ORF17	ORF18	ORF19	ORF20	ORF21

% H	89	66	98	87	66	66	66	7	1
F		6	65	B		-	-	-	-
Scor	558	503	580	373	1999	3044	1298		787
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatís						
£	L22180	M62819	L25105	L25105	125105	L25105	L25105		764n
Homology	ORF, 82 kDa protein	heat shock protein	GrpE-like protein	GrpE-like protein	has homology to putative heat shock proteins of Bacillus subtilis and Clostridium acetobutylicum; ORFA; putative	aminoacyl-tRNA synthetase	ORFB; putative	putative	المراهد المراعد المراهد المراهد المراهد المراهد المراهد المراهد المراهد المراع
stop	22153	22478	22824	23110	23394	24641	27710	27725	38740
begin	22602	22804	23183	23394	24569	26383	26640	28780	29957
ORF	ORF22	ORF23	ORF24	ORF25	ORF26	ORF27	ORF28	ORF29	Cr.730

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Score	200	, ,	192		663	681	959	755				892				142		462	801	154	355			350	
Species	pariling cubtiling	paciting subcilis	Escherichia coli		Mycobacterium tuberculosis	Chlamydia trachomatis	Chlamydia trachomatis	Flaveria bidentis				Pseudomonas aeruginosa				Nicotiana tabacum		Synechocystis sp.	Synechocystis sp.	Escherichia coli	Methanococcus Jannaschii			Pseudomonas syringae	DV. tabaci
ΩÏ	AFOORSOON	DE 000220	X67753		298209	L40822	L40822	L40958				M33223				L32794		D90911	X67516	M13045	U67500			U47017	
Homology	DAPH synthage-chorismate		arginine binding protein	putative	hypothetical protein MTCY154.05c	phophoglucoisomerase-like protein	phophoglucoisomerase-like protein	NADP-malate dehydrogenase	putative	putative	putative	membrane protein (arcD)	putative	putative	putative	dehydroquinate	dehydratase/shikimate dehydrogenase	3-dehydroquinate synthase	chorismate synthase	shikimate kinase II	5-enolpyruvylshikimate 3- phosphate synthase		putative	dihydrodipicolinate	דרים מר המשת
stop	39085		39927	40756	42007	43116	43802	45227	45275	45975	46506	46882	48178	48630	5005	50925		52348	53466	54595	55031	56084	58235	59181	
begin	38207		39151	39923	40760	42175	42999	44211	46072	46340	46895	47955	48585	2005	01/05	52439		53484	54536	55086	56350	55659	56847	58423	
ORF	ORF41		ORF42	ORF43	ORF44	ORF45	ORF46	ORF47	ORF48	ORF49	ORF50	ORF51	ORF52	ORF53	ORF54	ORFSS		ORF56	ORF57	ORF58	ORF59	ORF60	ORF61	ORF62	

ORF64         60188         61483         aspartokinase III           ORF65         61496         62353         dihydrodipicolinate           ORF66         62500         63141         putative           ORF67         63396         63983         hypothetical protein           ORF70         64944         64656         putative           ORF71         65347         67269         unknown           ORF72         67656         68873         putative           ORF74         69212         69721         high level kasgamycin           ORF75         69958         70455         polypeptide deformylase           ORF76         70701         71006         protein translocation           ORF77         73191         71006         protein translocation           ORF76         74876         homologous to unidentified           ORF87         74876         homologous to unidentified           ORF89         77124         75502         O530; This 530 a	A -	Species	Score	2
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69958 70455 polypeptide deformylase 70701 71006 protein translocation (secG) 73191 71086 putative 74900 73497 putative P. coli protein 75463 74876 homologous to unidentified P. coli protein 77124 75502 0530; This 530 aa ORF is 33 PCT identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HAEIN SW: P44808 77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog	D26185	Bacillus subtilis	306	43
70701 71006 protein translocation	Y10305	Calothrix PCC7601	272	43
Protein, low temperature (secG)	U32727	Haemonhilus influenzae	00	33
73191 71086 putative 74900 73497 putative 75463 74876 homologous to unidentified E. coli protein 77124 75502 0530; This 530 aa ORF is 33 pct identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HAEIN SW: P44808 77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog	ure		?	<del></del>
74900 73497 putative 75463 74876 homologous to unidentified E. coli protein 77124 75502 0530; This 530 aa ORF is 33 pct identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HAEIN 5W: P44808 77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog				
75463 74876 homologous to unidentified E. coli protein  77124 75502 0530; This 530 aa ORF is 33 pct identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HAEIN SW: P44808  77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog				
77124 75502 0530; This 530 aa ORF is 33 pct identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HAEIN SW: P44808 77000 77299 putative shows protein (xerC) 77145 integrase-recombinase protein (xerC) 78095 77145 hypothetical protein 81971 79878 LON protease homolog	M96343	Bacillus subtilis	283	8
pct identical (14 gaps) to 525 residues of an approx. 640 aa protein YHBS_HARIN SW: P44808   77000 77299 putative   protein (xerC)   78095 77145 integrase-recombinase   protein (xerC)   79065 78154 hypothetical protein   81971 79878 LON protease homolog	100000	- 1	!	]
525 residues of an approx. 640 aa protein YHES_HAEIN SW: P44808 77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog	aps) to	ina coli	1447	<del>7</del>
640 aa protein YHES_HAEIN SW: P44808 77000 77299 putative 78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog	approx.			
77000         77299         putative           78095         77145         integrase-recombinase           protein (xerC)         79065         78154         hypothetical protein           81971         79878         LON protease homolog	HAEIN			
78095 77145 integrase-recombinase protein (xerC) 79065 78154 hypothetical protein 81971 79878 LON protease homolog				
79065 78154 hypothetical protein 81971 79878 LON protease homolog	se U32750 Haemophilus	lus influenzae	495,	38
81971   79878 LON protease homolog	n D64001 Synechocystis	ystis sp.	400	40
	U88087	13	1927	48
ORF85 82639 83271 putative				

18	42						32	¥	35		53	36		0	4	42	97	37	80	?	51	48	88	
Score	822						128	671	120		542	326		487	2090	319	2411	707	177	1,1	634	558	300	
Species	Salmonella typhimurium						Bacillus subtilis	l oi	Bacillus subtilis		Bacillus subtilis	Helicobacter pylori		Helicobacter pylori	Thermus thermophilus	Helicobacter pylori	Chlamydia trachomatis	Sympohogyatia en	1.0		Haemophilus influenzae	Bacillus subtilis	Arabidopsis thaliana	
ti Ei	U58360						X12832	U12340	X17014		X06803	AE000583		AE000583	D49911	AE000583	U83196	D90903	D84432		U32723	L29475	U18969	
Homology	DnaJ	putative	putative	putative	putative	putative	Hpr protein	PTS enzyme I	ORF107	putative	dnaZX-like ORF put. DNA	1 70	10000 AD 00000 C	(uvrA)	UvrA	excinuclease ABC subunit A	pyruvate kinase	hypothetical protein	1	putative	exonuclease VII, large subunit (xseA)	triose phosphate isomerase	phosphoribosylanthranilate isomerase	putative
stop	84850	86921	87313	87805	88747	89265	89732	91447	91435	91745	92344	93361	94071	71011	94628	98113	98741	100337	101323	102210	102726	104254	105894	108460
begin	83792	84876	88650	87440	88400	41488	89355	89735	91749	92392	93138	94134	94637		98299	98715	100228	101347	102210	102485	104315	105075	105259	107429
ORF	ORF86	ORF87	ORF88	ORF89	ORF90	ORF91	ORF92	ORF93	ORF94	ORF95	ORF96	ORF97	ORF98		ORF99	ORF100	ORF101	ORF102	ORF103	ORF104	ORF105	ORF106	ORF107	ORF108

2	,			100		۶/		50	3 2	38	62	91	20		S.	3 4		T	T	45	?	84	2	2	5 8	3 8	3	51
Score				2007		313		443	52.8 52.8	143	254	2675	3486	)	677	121				1062	3	790		188	2	000	3	1763
Species	4			Chlamydia trachomatis	Homomby 1 ! A. E.	naemophilus iniluenzae		Bacillus subtilis			Synechocystis PCC6803	Staphylococcus aureus	Haemophilus influenzae	•	Homo saniens		1			Gallus gallus		Haloferax volcanii		Enterococcus hirae				Synechocystis sp.
q				L22216	1132754	#C / 7CO		D13303	Z11839	211839	X53178	X64172	U32733		L19437	U67611				U22077		X79516		D17462	X76913	X76913		D64006
Homology	putative	putative	putative	elongation factor Tu	transcription	antitermination protein	(Ssnu)	ribosomal protein L11	ribosomal protein L1	ribosomal protein L10	rpl12 (AA 1-128)	DNA-directed RNA polymerase beta chain	DNA-directed RNA polymerase	beta' chain (rpoC)	transaldolase	transaldolase	putative	putative	putative	Al isoform of vacuolar H+-	ATPase subunit A	membrane ATPase	putative	Na+ -ATPase subunit I	v-type Na-ATPase	v-type Na-ATPase	putative	valyl-tRNA synthetase
stop	108955	109013	109704	112520	113463			113994	114604	115253	115676	119795	124010		124988	125106	125536	126930	127785	129714		131033	131629	133156	133584	133999	134508	137454
begin	108665	109459	110366	111330	112915			113566	114020	114720	115362	116022	119823		124065	124873	126261	126328	127138	127924		129720	131018	131834	133075	133625	133861	134638
ORF	ORF109	ORF110	ORF111	ORF112	ORF113			ORF114	ORF115	ORF116	ORF117	ORF118	ORF119		ORF120	ORF121	ORF122	ORF123	ORF124	ORF125		ORF126	ORF127	ORF128	ORF129	ORF130	ORF131	ORF132

ORF	beqin	Stop	Homology	4			
20.000	1226	3,000,		77	Species	Score	æ H
OKFISS	13/442	140276	PknD	Z95209	Mycobacterium	452	44
					tuberculosis		
ORF134	140733	140335	putative				
ORF135	141799	141077	porphobilinogen deaminase	U22968	Yersinia pestis	282	38
ORF136	143240	141780	unknown	D26185	Bacillus subtilis	1113	53
ORF137	143829	143128	ORF3	D64116	I	356	39
ORF138	143923	144393	putative				
ORF139	144548	146326	unknown	247210	Streptococcus pneumoniae	741	44
ORF140	146413	147078	manganese superoxide dismutase precursor	D12984	Caenorhabditis elegans	625	56
ORF141	147140	148075	acetyl-CoA carboxylase beta subunit (accD)	AE000604	Helicobacter pylori	704	52
ORF142	148115	148549	Dut	296072	Mycobacterium tuberculosis	277	53
ORF143	148524	149027	enzyme IIANtr	U18997	Escherichia coli	168	44
ORF144	149000	149305	putative		1		
ORF145	149187	149708	enzyme IIANtr	U18997	Escherichia coli	169	43
ORF146	149712	150911	putative		ı		
ORF147	152044	151004	putative				
ORF148	152664	151999	putative				
ORF149	152900	153352	hypothetical	U32702	Haemophilus influenzae	292	47
ORF150	153389	153997	hypothetical protein in purB 5' region	AE000213	Escherichia coli	555	49
ORF151	155276	153984	ClpC adenosine triphosphatase	U02604	Bacillus subtilis	986	45
ORF152	156544	155231	ClpC adenosine triphosphatase	U02604	Bacillus subtilis	1535	53
ORF153	156806	157525	putative				
ORF154	157489	158955	Unknown	Y08559	Bacillus subtilis	, 66	39
ORF155	159104	159961	putative		ı		
ORF156	159916	161220	putative				

1%	35	51	55	2	45	33	42		48		47	44	43	14	38		20	14	
Score	175	524	543	287	449	111	722		462		210	176	789	177	110		558	453	
Species	Saccharomyces cerevisiae	Vibrio alginolyticus	Vibrio alginolyticus	Haemophilus influenzae	Vibrio alginolyticus	Helicobacter pylori	Bacillus subtilis		Synechocystis sp.		Neisseria flavescens	Neisseria flavescens	Bacillus subtilis	Helicobacter pylori	Pseudomonas aeruginosa		Synechococcus PCC7942	Haemophilus influenzae	
a	U12980	D49364	237111	U32702	Z37111	AE000652	X02369		D90906		M26645	M26645	215056	AE000589	L35259		U59236	U32691	
Homology	glycine cleavage protein homolog	unidentified protein of Na+- translocating NADH-quinone reductase	NADH:uniquinone oxidoreductase	NADH:ubiquinone oxidoreductase (GP:Z37111_4)	NADH:ubiquinone oxidoreductase subunit B	H. pylori predicted coding region HP1542	pot. ORF 446 (aa 1-446)	putative	hypothetical protein	putative	penicillin-binding protein 2	penicillin-binding protein 2	murE gene product	N-acetylmuramoyl-L-alanine amidase (amiA)	integration host factor beta subunit	putative	carboxyltransferase alpha subunit	ATP dependent translocator homolog (msbA)	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1
stop	161593	161623	162363	162994	164474	166093	166729	168848	170431	171334	172376	172722	174496	174968	175710	177009	178115	180021	180048
begin	161183	162354	163013	163941	165505	166686	168171	169249	169586	170780	171333	172309	173048	174399	175267	175714	177423	178084	180704
ORF	ORF157	ORF158	ORF159	ORF160	ORF161	ORF162	ORF163	ORF164	ORF165	ORF166	ORF167	ORF168	ORF169	ORF170	ORF171	ORF172	ORF173	ORF174	ORF175

1%	¥	S	53	43	30	35		36		38				9	43		56	53	42	20
Score	256	173	371	452	93	154		96		66				373	545		1758	580	148	795
Species	Helicobacter pylori	Arabidopsis thaliana	Bacillus subtilis	Bacillus subtilis	Escherichia coli	Haemophilus influenzae		Synechocystis sp.		Rhodobacter sphaeroides				Helicobacter pylori	Staphylococcus aureus		Homo sapiens	Escherichia coli	Saccharomyces cerevisiae	Caenorhabditis elegans
A	AE000536	AF007270	Y13937	AF008220	D90888	U32728		D90902		AJ000977				AE000645	D89066		U47025	X16931	X86470	277659
Homology	H. pylori predicted coding region HP0152	contains similarity to DNA polymerase III, alpha chain (SP:P47277)	putative Ptcl protein	Nifs2	similar to [SwissProt Accession Number P37908]	hypothetical	putative	regulatory protein for beta- lactamase	putative	prolipoprotein diacylglyceryl transferase	putative	putative	putative	60 kDa inner-membrane protein	DnaA	putative	glycogen phosphorylase B	glycogen phosphorylase (AA 1 - 790)	unknown	F23B12.5
stop	180631	181398	183656	184786	184796	186000	186749	187809	188798	190352	190510	191786	192464	193183	194630	194690	197031	197635	198208	197668
begin	181398	182594	182895	183665	186007	186848	187270	187426	189481	189693	190235	190785	191790	192392	193254	195046	195184	197018	197762	198963
ORF	ORF176	ORF177	ORF178	ORF179	ORF180	ORF181	ORF182	ORF183	ORF184	ORF185	ORF186	ORF187	ORF188	ORF189	ORF190	ORF191	ORF192	ORF193	ORF194	ORF195

ORF	begin	aton	Homology	1			
70,000	11600	2000	Yearon	מד	Species	Score	1%
OKF 196	199957	198962	pyruvate dehydrogenase El beta subunit	U09137	Arabidopsis thaliana	856	48
ORF197	200327	199941	pyruvate dehydrogenase El component, alpha subunit	U38804	Porphyra purpurea	170	31
ORF198	200685	200266	pyruvate dehydrogenase complex El alpha subunit	U81808	Thiobacillus ferrooxidans	302	9
ORF199	200962	200585	TPP-dependent acetoin dehydrogenase alpha-subunit	L31844	Clostridium magnum	127	43
ORF200	201169	202377	putative				
ORF201	203441	202380	UDP-3-0-[3-hydroxymyristoyl]	U70214	Escherichia coli	577	38
			glucosamine N- acyltransferase				
ORF202	203998	203471	putative				
ORF203	206449	204059	OMP1 precursor	U51683	Brucella abortus	83	31
ORF204	207425	206811	recombination protein	D90916	Synechocystis sp.	334	40
ORF205	207506	208528	beta-ketoacyl-acyl carrier protein synthase III	M77744	Escherichia coli	206	20
ORF206	208545	209471	malonyl-CoA:Acyl carrier protein transacylase	U59433	Bacillus subtilis	522	48
ORF207	209471	210214	3-ketoacyl-acyl carrier protein reductase	U59433	Bacillus subtilis	616	51
ORF208	210586	210816	acyl carrier protein (acpP)	U32701	Haemophilus influenzae	220	22
ORF209	211332	210883	protein kinase type II regulatory subunit (, EC	J02934	Rattus norvegicus	150	31
			2.7.1.37)				
ORF210	212978	211374	putative				
ORF211	214134	212875	unknown	AF017105	Chlamydia psittaci	852	83
ORF212	214710	214168	inclusion membrane protein C	AF017105	Chlamydia psittaci	231	43

Г				1		$\overline{}$	-		· -	_	$\neg$	$\overline{}$			_									
•	47	0/	S	3									66	46	40	25	42	36		48	35	8		35
1	181	1341	1001	7707									619	000	250	334	96	321		274	246	0	•	740
Sportos	Chlamydia psittaci	Chlamydia psittaci	Chlamidia noittani	- 1									Chlamydia trachomatis	Bacillus subtilie	Education Substates	Methanococcus jannaschii	Saccharomyces cerevisiae	Bacillus firmus		Helicobacter pylori	Escherichia coli			Lactococcus lactis
QI	AF017105	AF017105	AF017105										US7090	D84432		U67467	U43834	U61168		AE000602	AE000232			AF005098
Homology	inclusion membrane protein B	sodium-dependent transporter	amino acid transporter	putative	putative	putative	putative	putative	putative	putative	putative	putative	LAGLI-DADG endonuclease	YqfU	-1 com.	phenylacryllc acid decarboxylase	Ydr537cp	4-hydroxybenzoate	putative	stationary-phase survival protein (surE)	f311; This 311 aa ORF is 22	pct identical (13 gaps) to 186 residues of an approx.	SW: P37563; pyul of D21139	GadC
stop	214754	215236	216892	217441	218702	219009	219748	220430	221074	221541	222092	223290	223818	225171	225174	£/1C22	225549	225749	226769	227161	227750			228607
begin	215143	216705	217917	217088	218364	218695	219179	219891	220499	221137	221601	222472	223423	224278	225749		225334	226654	227299	227646	228457		· · · · · · · · · · · · · · · · · · ·	230001
ORF	ORF213	ORF214	ORF215	ORF216	ORF217	ORF218	ORF219	ORF220	ORF221	ORF222	ORF223	ORF224	ORF225	ORF226	ORF227		ORF228	ORF229	ORF230	ORF231	ORF232			ORF233

ORF	begin	stop	Homology	£	Chanter	1	5
166400	221024	1 1 10 10		7	sarnaďe	SCOL	T.%
OKF234	\$/OT67	230151	1; Inis 374 aa ORF	AE000299	Escherichia coli	985	65
			identical (9				
			102 residues of an approx.				
			512 aa protein FLIC_SALMU			<u></u>	
00000	221240	00000					
OKF 235	231348	233006	putative				
ORF236	233059	233829	orf2	D88555	Methanobacterium	351	52
2000	.000		- 1		thermoautotrophicum		
OKF 23 /	73380T	234265	hypothetical protein	D90906	Synechocystis sp.	151	37
ORF238	234282	234854	ORF_0211	U28377	Escherichia coli	105	Ŋ
ORF239	236300	235227	glutamate 1-semialdehyde 2,1	X82072	Pseudomonas aeruginosa	650	52
			aminomutase		)	-	
ORF240	236314	238209	leucine tRNA synthetase	AF008220	Bacillus subtilis	1836	61
ORF241	238164	238769	leucine tRNA synthetase	AF008220	Bacillus subtilis	410	46
ORF242	238769	240061	3-deoxy-D-manno-2-	222659	Chlamydia trachomatis	2240	100
			octulosonic acid (Kdo)			)     	
			transferase				
ORF243	242022	240313	pyrophosphate-dependent	232850	Ricinus communis	1021	43
			phosphofructokinase beta				
			subunt				
ORF244	242846	241941	putative				
ORF245	244480	242798	pyrophosphate-dependent	232850	Ricinus communis	1017	42
			phosphofructokinase beta				
			subunit				
ORF246	245897	244479	YflS	D86417	Bacillus subtilis	951	42
ORF247	246877	245924	putative				
ORF248	247731	246985	ATP binding protein	L18760	Lactococcus lactis	442	47
ORF249	248585	247743	sporulation protein	M57689	Bacillus subtilis	532	38
ORF250	249420	248569	sporulation protein	M57689	Bacillus subtilis	601	88
ORF251	250383	249766	sporulation protein	M57689	Bacillus subtilis	464	47
ORF252	251186	250545	oligopeptide permease homolog AII	AF000366	Borrelia burgdorferi	119	31
ORF253	252111	251095	sporulation protein	M57689	Bacillus subtilis	317	36
					-1		;

P36175 (660)  """  """  """  """  """  """  """						Synechocystis sp.  Zymomonas mobilis Salmonella typhimuriu Aeromonas salmonicida Bacillus subtilis Haemophilus influenza Anabaena sp. Chlamydia trachomatis	Synechocystis sp. Zymomonas mobilis Salmonella typhimuriu Aeromonas salmonicida Bacillus subtilis Haemophilus influenza Anabaena sp. Chlamydia trachomatis Chlamydia trachomatis	Synechocystis sp. Zymomonas mobilis Salmonella typhimuriu Aeromonas salmonicida Bacillus subtilis Haemophilus influenza Anabaena sp. Chlamydia trachomatis Chlamydia trachomatis	Synechocystis sp.  Zymomonas mobilis  Salmonella typhimuriu  Aeromonas salmonicida  Bacillus subtilis  Haemophilus influenza  Anabaena sp.  Chlamydia trachomatis  Chlamydia trachomatis  Chlamydia trachomatis  Chlamydia trachomatis
	D90905 L33777 X07916	0905 3777 7716 7916	905	05 77 77 16 16 78 85					<del></del>
D909	D909	909	5		55 . 00 . 2	005 777 778 885 002 002 95	05 77 77 16 16 02 02 02 53 53	05 77 78 85 85 02 02 95 95	05 77 77 16 16 02 02 02 02 95 95
1 1 1	<del></del>	T X X T T T T T T T T T T T T T T T T T	X07 X07 L147 D26	Y079 Y079 L479 U328					
Mg2+ transporter tRNA guanine transglycosylase putative putative	Mg2+ transporter tRNA guanine transglycosylase putative putative subunit B of DNA gyrase	trans guani guani ive iive nit B	trans guanj guanj sglycc live lit B lit B	guani guani gglycc ive iive iit B iit B	transpo guanine sglycosy ive iive of somenta in hete	transporter guanine sglycosylase live live lit B of DNA gyr lyrase lyrase live isozyme of lehydrogenase; opmentally regu in heterocyst opment se 6-phosphate rogenase	transporter guanine sglycosylase live live lit B of DNA gyr lit B of DNA gyr live isozyme of ehydrogenase; opmentally regu in heterocyst opment se 6-phosphate rogenase se 6-phosphate	transporter guanine seglycosylase tive tive nit B of DNA gyr gyrase own ication protein tive isozyme of dehydrogenase; lopmentally regu in heterocyst lopment ose 6-phosphate drogenase drogenase	transporter guanine silve live live lit B of DNA gyr live isozyme of lehydrogenase; opmentally regu in heterocyst opment coment se 6-phosphate rogenase se 6-phosphate rogenase
transglycc putative putative	transglycc transglycc putative putative subunit B	transglycc transglycc putative putative subunit B	transglycc putative putative putative subunit B DNA gyrase	tRNA guanine transglycosy putative putative subunit B of DNA gyrase unknown replication	transglycosy putative putative putative subunit B of DNA gyrase unknown replication replication putative iso 6-P-dehydrog developmenta gene in hete development	transglycosy  putative  putative  subunit B of  nuknown  replication  replication  putative iso  6-P-dehydrog  developmenta  gene in hete  development  gene in hete  development  gene in hete  development  gene in hete  development	transglycosy  putative  putative  subunit B of  nuknown  replication  putative iso  6-P-dehydrog  developmenta  gene in hete  development  gene in hete  development  glucose 6-ph  dehydrogenas  glucose 6-ph  dehydrogenas	transglycosy putative putative subunit B of nuknown replication replication ce-P-dehydrog developmenta gene in hete development glucose 6-ph dehydrogenas glucose 6-ph dehydrogenas ORF3	transglycosy putative putative putative subunit B of nuknown replication putative iso 6-P-dehydrog developmenta gene in hete development gene in hete development gene of ph dehydrogenas glucose 6-ph dehydrogenas ORF3 ORF3
putative putative	putative putative subunit B	putative putative subunit B DNA gyrase	putative putative putative subunit B DNA gyrase	putative putative subunit B DNA gyrase unknown replicatio					
-									
	subunit B	subunit B	Subunit B DNA gyrase	Subunit B of DNA gyrase unknown replication	DNA gyrase  Unknown  replication  putative isc  6-P-dehydrog  developmenta  gene in hete  development	DNA gyrase  unknown  replication  putative iso  6-P-dehydrog  developmenta  gene in hete  development  gene in hete  development	DNA gyrase  unknown  replication  putative iso  6-P-dehydrog  developmenta  gene in hete  development  glucose 6-ph  dehydrogenas  glucose 6-ph  dehydrogenas	DNA gyrase  unknown  replication  putative iso  6-P-dehydrog  developmenta  gene in hete  development  glucose 6-ph  dehydrogenas  glucose 6-ph  dehydrogenas	DNA gyrase  unknown replication putative iso 6-P-dehydrog developmenta gene in hete development gene on hete development gene of ph dehydrogenas glucose 6-ph dehydrogenas ORF3

*		40			41	04	37	52			T	98	37	Ţ	45	41	28			$\top$
Score		82			745	197	292	316				123	200	+	169	147	167			-
Species		Escherichia coli			Thermus aquaticus	Escherichia coli	Mycoplasma pneumoniae	Mycobacterium tuberculosis				Escherichia coli	Bacillus subtilia		Klebsiella pneumoniae	Bacillus subtilis	Spirulina platensis			
QI		U73857			M74792	D90870	AE000047	Z84395				D90832	M57689		X58778	Z38002	S70419			
Homology	putative	similar to putative oxygenase of S. fradiae	putative	putative	DNA ligase	DNA LIGASE (EC 6.5.1.2) (POLYDEOXYRIBONUCLEOTIDE SYNTHASE (NAD+)).	Mycoplasma pneumoniae, DNA ligase; similar to Swiss- Prot Accession Number P15042, from E. coli	unknown	putative	putative	putative	Preprotein translocase SecA subunit.	sporulation protein	putative	orfX gene product	Similar to Saccharomyces cerevisiae SUA5 protein	serine esterase (Spirulina platensis, C1, Peptide, 207 aa)	putative	putative	putative
Btop	298735	300458	300527	302071	304973	306111	306436	306977	309276	309872	310716	311972	314364	314738	314741	315665	316529	317338	317499	317874
begin	297791	298905	302152	304917	306157	306494	306963	308773	309881	310720	311570	312451	313435	314340	315526	316507	317284	317592	318470	317599
ORF	ORF296	ORF297	ORF298	ORF299	ORF300	ORF301	ORF302	ORF303	ORF304	ORF305	ORF306	ORF307	ORF308	ORF309	ORF310	ORF311	ORF312	ORF313	ORF314	ORF315

ORF	begin	stop	Homology	1	- 7 7		
ORF316	318947	318477	putative	3	Species	Score	¥
ORF317	319342	320142	ORF2	L35036	Chlamydia neittari	200	G.
ORF318	320544	321497	putative		- 1	200	3
ORF319	321485	321937	putative				I
ORF320	321901	322362	putative				
ORF321	322301	323140	putative				
ORF322	323144	324913	putative				
ORF323	325621	324977	Yqiz	D84432	Bacillus subtilis	430	43
ORF324	326268	325621	integral membrane protein homolog	U97348	Lactobacillus fermentum	343	4
ORF325	326469	327203	adenylate kinase	AB000111	Synechococcus sn	271	98
ORF326	327281	328150	putative			7/5	ş
ORF327	328605	328204	RpsI	295389	Mycobacterium	315	55
ODE328	330066	220224	- [		ruperculosis		
ONE 320	323000	328/34	508 ribosomal subunit protein L13	U18997	Escherichia coli	269	09
ORF329	329663	329292	хчъх	D84432	Bacillus subtilis	207	£
ORF330	330666	329608	biotin carboxylase	L14862	J	1080	3 8
ORF331	331161	330670	хдъм	D84432	1	5001	3 6
ORF332	331731	331177	elongation factor P	D64001		200	35
ORF333	332404	331721	putative CfxE protein	X13937	Barillus cubtilis	167	3 2
ORF334	332779	333021			- 1	202	3
ORF335	333005	333589	putative				
ORF336	334357	333806	putative				
ORF337	334089	334361	putative				
ORF338	335142	334729	putative				T
ORF339	335195	335602	putative				T
ORF340	335673	335194	putative				
ORF341	336334	335903	putative				
ORF342	337378	336338	putative			1	
ORF343	339947	337347	ATP-dependent protease binding subunit	M29364	Escherichia coli	2005	53
						-	-

1%	39	48	66	66	100	100	æ	41	46	4		2	42	
Score	208	140	361	1271	1051	344	344	387	492	397		606	113	
Species	Bacillus licheniformis	Streptococcus agalactiae	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia psittaci	Escherichia coli	Homo sapiens	Escherichia coli		Brassica napus	Synechocystis sp.	
a	D88209	U49821	M31739	L12004	M31739	M31739	S40172	AE000174	U63329	AE000209		\$60064	D90914	
Homology	Pz-peptidase	group B oligopeptidase PepB	hypA protein	heat shock protein	hypB protein	hypB protein	orf 3'of chaperonin homolog hypB [Chlamydia psittaci, pigeon strain P-1041, Peptide Partial, 98 aa]	o247; This 247 aa ORF is 51 pct identical (0 gaps) to 117 residues of an approx. 160 aa protein YPH7_CHRVI SW: P45371	mutY homolog	hypothetical 36.0 kD protein in rne-rpmF intergenic region	putative	enoyl-acyl carrier protein reductase [Brassica napus, Peptide, 385 aa]	hypothetical protein	putative
stop	341847	342022	342470	343370	344032	344225	345142	345161	346080	347940	348146	351283	351314	352245
begin	340507	341783	342249	342597	343361	343956	344357	345934	347102	347113	350164	350423	352207	352727
ORF	ORF344	ORF345	ORF346	ORF347	ORF348	ORF349	ORF350	ORF351	ORF352	ORF353	ORF354	ORF355	ORF356	ORF357

ORF	begin	stop	Homology	d.	Chantan		j
ORF358	353709	353305	ETINCTION INENOGEN CIMILAD	200.000	4!	SCORE	*
			IN E. COLI A PNEUMON	AB001488	Bacillus subtilis	213	9
ORF359	354218	353670	NADPH thioredoxin reductase	223108	Arabidopsis thaliana	577	99
ORF360	354721	354140	Thioredoxin Reductase (NADPH)	D45049	Neurospora crassa	417	99
ORF361	354966	356672	30S ribosomal protein S1	D90729	Escherichia coli	1305	44
ORF362	356700	357377	NusA	U74759		948	100
ORF363	357326	358093	NusA	U74759	Chlamydia trachomatis	1216	100
ORF364	358035	360743		U74759	Chlamydia trachomatis	3311	86
ORF365	360753	361121	ORF6 gene product	218631	Bacillus subtilis	116	33
ORF366	361162	361884	tRNA pseudouridine 55 synthase	D90917		362	45
ORF367	361826	362746	protein X	M35367	Pseudomonas fluorescens	192	49
ORF368	363859	362816	hypothetical GTP-binding protein in pth 3' region	AE000219	Escherichia coli	978	25
ORF369	364116	365195	cds1 gene product	U88070	Chlamydia psittaci	1631	88
ORF370	365198	365587	cds2 gene product	U88070		516	93
ORF371	365479	367320	gene	U88070	Chlamydia psittaci	2817	87
ORF372	367341	368603	gene	U88070	Chlamydia psittaci	585	37
OKF 3 / 3	368644	369081	luct	U88070	Chlamydia psittaci	528	29
OKF3/4	369088	370251		U88070	Chlamydia psittaci	1362	62
6/64	60/0/6	3/1086	ribosomal protein L28 (rpL28)	U32776	Haemophilus influenzae	182	46
ORF376	371203	372816	hypothetical protein	U88070	Chlamydia psittaci	1926	89
ORF377	373119	373529	hypothetical protein	U88070	1	286	64
ORF378	373614	374204	hypothetical protein	U88070		379	48
ORF379	374736	374224	putative				

	stop	Homology	a	an journal	2	ļ
374703	┢	putative		200	Score	P.1
376748 c	0 6	corresponds to a 97 amino acid long polypeptide	L40838	Chlamydia trachomatis	490	86
378737 me	me de	methylenetetrahydrofolate dehydrogenase	D64000	Synechocystis sp.	678	21
	Z.	putative				
379403 hy	ьу	hypothetical	U32702	Haemophilus influenzae	137	45
379641 sm	Sm	small protein	D90914	Cracochoods		-
381470 DN	NO Bu	DNA polymerase III beta- subunit (dnaN)	U32780	Haemophilus influenzae	917	39
382567 re	re	recombination protein	D26185	Bacillus subtilis	7.7.7	36
	ρα	putative			//#	S
	Кq	hypothetical	U70214	Escherichia coli	124	35
383949 put	hnd	putative			101	3
385178 cons	con	conserved hypothetical secreted protein	AE000606	Helicobacter pylori	185	45
	hypc	hypothetical protein	D64000	Synechocystis sp	282	41
	puta	putative		ł		-
386872 ORF	ORF	ORF1; putative	M26130	Streptococcus	150	35
387338 ytgD	ytgi	0	AF008220	Barillus subtilia	90,	Ş
387816 TroR	Trol	8	U55214	Treponema nallidum	134	7,
388237 put	put	1	U30821		164	31
380173 mg	am F	amino acids				
+	71	)B	U55214	Treponema pallidum	592	51
389945 YtgA	Ķ	ЭA	AF008220	Bacillus subtilis	282	30
-	Put	putative		1		
393996 ade	ade	adenine nucleotide	Z49227	Arabidopsis thaliana	1295	5,6
	tr	translocase			)	3
-	le]	lepA gene product	X91655	Bacillus subtilis	1235	9
395992 GTI	E :	GTP-binding membrane protein (lena)	AE000552	Helicobacter pylori	543	28
		(Trial)				

*I	55	51	45	41	35	39	22		49		43	1	;	•	SS.		4							36		
Score	411	806	844	463	134	355	98		995		166	622			1537		496							172		
Species	Homo sapiens	Candida albicans	Bacillus caldotenax	Streptoverticillium griseocarneum	Bacillus subtilis	Caulobacter crescentus	Rhodobacter capsulatus		Synechocystis sp.		Escherichia coli	1.5			Rhodobacter capsulatus		Bacillus subtilis							Mycobacterium	2100112	
a	U30255	AB006102	M13148	X68709	X63698	M73782	M59855		D90908		U50134	M27141			U41762	00.400	D64432							294752		
Homology	phosphogluconate dehydrogenase	6-phosphogluconate dehydrogenase	tyrosyl-tRNA synthetase	whiG-Stv gene product	FLHA gene product	flbF	ferredoxin IV	putative	gdog	putative	YfiH	dihydrolipoamide	transsuccinylase (odhB; EC	2.3.1.61)	alpha-ketoglutarate	VeeD	, 10 ht	pucacive	pararra	pucative	putative	putative	putative	unknown	putative	putative
stop	396059	396542	397401	398909	399778	400034	402021	403220	405180	403276	405920	405955			407056	411416	01717	413410	2000	413732	415112	413978	415177	416740	417721	418031
begin	396538	397507	398753	399688	400167	401224	401776	402126	403348	403788	405165	407049			409773	410532	411707	413433	412404	F0F57#	413841	414379	416664	417456	418053	418603
ORF	ORF404	ORF405	ORF406	ORF407	ORF408	ORF409	ORF410	ORF411	ORF412	ORF413	ORF414	ORF415			ORF416	ORF417	ODE418	ORF419	00000	OVETE	ORF421	ORF422	ORF423	ORF424	ORF425	ORF426

4.	92	96	41	99	3	55		53	8	}		æ				53	¥	T	47	47	56	57
Score	1661	612	269	165		1229		633	612			173				289	817		395	287	374 .	199
Species	Chlamydia trachomatis	Chlamydia trachomatis	Mycoplasma salivarium	Mycobacterium leprae		Synechocystis sp.		Pseudomonas aeruginosa	Helicobacter pylori			Saccharomyces cerevisiae				Escherichia coli	Schizosaccharomyces pombe		Lactococcus lactis	Synechocystis sp.	12	Haemophilus influenzae
A	L10193	M86605	D17450	L39923		D90908		D83138	AE000554			U27182				U29581	AB004537		AF005098	X72627	U32705	U32705
Homology	Hc2 nucleoprotein	[karp] gene products	aminopeptidase	putative	putative	glycogen operon protein GlgX	putative	Holliday junction specific DNA helicase	deoxycytidine triphosphate	e (dcd	putative	biotin apo-protein ligase	putative	putative	putative	exonuclease V alpha-subunit	methionyl-tRNA synthetase	putative	RNAseH II	ribosomal protein L19	tRNA (guanine-N1) - methyltransferase (trmD)	tRNA (guanine-N1)- methyltransferase (trmD)
stop	418647	419672	420245	421518	423043	425079	425146	426245	427817		429886	429857	430323	431787	431987	434475	434620	436272	436567	437894	438285	438986
begin	419531	420190	421171	421988	422486	423226	426054	426985	427248		429560	430360	430637	430933	431658	432232	436308	436574	437685	438262	439127	439339
ORF	ORF427	ORF428	ORF429	ORF430	ORF431	ORF432	ORF433	ORF434	ORF435		ORF436	ORF437	ORF438	ORF439	ORF440	ORF441	ORF442	ORF443	ORF444	ORF445	ORF446	ORF447

	pegin	stop	Homology	αI	Sheries	0000	ð
ORF448	439705	439358	ribosomal protein S16 (rpS16)	U32705	Haemophilus influenzae	168	39
ORF449	441042	439699	signal recognition particle protein	AE000347	Escherichia coli	865	40
ORF450	441911	441042	product similar to B.coli PRFA2 protein	249782	Bacillus subtilis	314	37
ORF451	442593	441898	polypeptide chain release factor 1 (prfA)	U32830	Haemophilus influenzae	708	62
ORF452	444505	446388	leader peptidase I	D90904	Synechocystis sn	268	44
ORF453	448068	446452	isoleucyl-tRNA synthetase	U04953	1	200	1 0
ORF454	449575	447932	isoleucyl-tRNA synthetase	U04953		1697	55
ORF455	450546	451076	putative			1001	3
ORF456	451623	451144	putative				
ORF457	452593	451517	putative				
ORF458	453195	452632	putative				
ORF459	453567	454868	product similar to E. coli PhoH protein	297025	Bacillus subtilis	820	20
ORF460	455430	454972	CydB	295554	Mycobacterium tuberculosis	105	31
ORF461	456047	455367	cyanide insensitive terminal oxidase	Y10528	Pseudomonas aeruginosa	388	88
ORF462	457384	456047	cyanide insensitive terminal oxidase	Y10528	Pseudomonas aeruginosa	537	25
ORF463	457659	458450	YbbP	AB002150	Bacillus subtilis	700	12
ORF464	458508	459632	putative		- 1	1,20	1
ORF465	459839	461203	HtrB protein	X61000	Escherichia coli	77	2
ORF466	461624	461196	unknown	U87792		114	5 8
ORF467	461887	462621	hypothetical protein	275208	Bacillus subtilis	148	3 5
ORF468	463758	462895	putative				; T
ORF469	464048	464629	putative				
ORF470	464721	465848	putative				
ORF471	467420	466113	PBT112	D90913	Synechocystis sp.	892	48

*	46					ಜ		39	88		g	9 04		35	40	 :		S.	3	T	ç	2 2	<u> </u>	39	3				T	25
Score	1051					173		175	193		100	537		234	313	}		191	1		114	777	:	506	)			-		1624
Species	Moraxella catarrhalis					Chlamydia psittaci	- 1	Chlamydia psittaci	Chlamydia psittaci		Chlamydia psittaci			Chlamydia psittaci	Chlamydia psittaci			Chlamydia psittaci			Chlamydia psittaci	1		Chlamydia psittaci						Synechococcus PCC6301
£	U49269					U72499		U65942	U72499		U65942	U72499		U72499	U72499			U72499			U65942	U72499		U72499						M31544
Homology	amidase	putative	putative	putative	putative	putative 98 kDa outer	DOMPSON TESTINE	r 3 . I	putative 98 kDa outer membrane protein	putative	POMP91A	putative 98 kDa outer membrane profein	Ton Table	purative 98 kba outer membrane protein	putative outer membrane	protein	putative	putative 98 kDa outer	membrane protein	putative	POMP90A precursor	putative 98 kDa outer	membrane protein	putative 98 kDa outer	membrane protein	putative	putative	putative	putative	branching enzyme
stop	467419	468906	469675	469826	471106	473267	473695	213030	474527	474602	475613	476517	47866	C000/#	479088		479668	479895		481496	483429	484964		487864		485222	489247	488233	490456	490507
begin	468891	469280	469349	471226	471624	471954	473252	202012	4/3982	475198	476527	478640	479084	F0000	479723		480012	481466		481732	481864	483402		484898		485725	488204	488571	489440	492765
ORF	ORF472	ORF473	ORF474	ORF475	ORF476	ORF477	ORF478	OPEA70	OKF 4 / 9	ORF480	ORF481	ORF482	ORF483		ORF484		ORF485	ORF486		ORF487	ORF488	ORF489		ORF490		ORF491	ORF492	ORF493	ORF494	ORF495

		Homology	Ωī	Species	Score	1%
492893 putative	ati	ive				
492737 putative	1.2	ive				
	. /		D84432	Bacillus subtilis	230	4
			M54884	Escherichia coli	245	84
495365 putative	٠.	ve				
494872 putative	_	/e				
496634 putative		ve				
497176 putative		ve				
498515 putative		ve				
-		ve				
-	-	ye.				
500790 penicil (1ytB)	[편	penicillin tolerance protein (lyt8)	U32781	Haemophilus influenzae	702	50
501808 putative	>	U				
502692 putative	>	U				
503722 hypothetical	er er	ical protein	296072	Mycobacterium	102	42
506986 hypotherical	i a	protein in	0100004			
prs	te	ic region		Escherichia coll	740	<del>1</del>
$\dashv$	5	dì				
	اند	fumarate hydratase	AF013216	Myxococcus xanthus	611	\$
~	>	Ð				
	י מט	e	D64000	Synechocystis sp.	386	57
510691 thiamin		thiamine-repressed protein	U32720	Haemophilus influenzae	82	31
	- 1				_	
511527 putative	<b>&gt;</b>	ð				
512185 hypothetical	1	ical protein	U67608	Methanococcus	208	39
(SP:P46851)	36	151)		jannaschii	)	}
513092 methionine	ď	ine amino peptidase	M15106	Escherichia coli	384	46
513055 putative		a			-	
╁		Ze Ze				
515269 putative		ē				
	1		7			

19	9			51	52	49	47	\$	40	9/		44	•						52	40	47	SS.	43
Q TO DE		-		340	245	306	387	860	070	314		148	0.54						476	164	230	452,	488
Species				Porphyromonas gingivalis	Synechocystis sp.	1	Spinacia oleracea	Escherichia coli	Escherichia coli			Porphyra purpurea	mand 1 1 1						Bacillus subtilis	Bacillus subtilis	Bacillus stearothermonhilus	Mycobacterium tuberculosis	Synechococcus PCC7942
A				X95938	D90901	AF010578	U52048	U18997	U01376	U32812		U38804							X53057	275208	X16188	285982	U86147
Homology	putative	putative	putative	orf150 gene product	30S ribosomal protein S15	polynucleotide phosphorylase	polyribonucleotide phophorylase	polynucleotide phosphorylase	ATP-binding protein	cell division protein (ftsH)	putative	ORF327 gene product	putative	putative	putative	putative	putative	putative	phenylalanyl-tRNA synthetase alpha subunit	phenylalany-tRNA synthetase beta subunit	ribosomal protein L20 (AA 1-119)	unknown	UDP-N- acetylenolpyruvylglucosamine reductase
stop	515804	516422	517171	517400	518380	518822	519516	520497	520718	521886	522143	523623	525746	526078	526400	526735	526851	528292	529142	529624	530223	530737	533272
begin	515553	515808	516476	517927	518096	518403	518923	519577	521986	522131	523495	524591	524652	525731	525939	526301	528323	528861	529723	530166	530543	531378	532370
ORF	ORF523	ORF524	ORF525	ORF526	ORF527	ORF528	ORF529	ORF530	ORF531	ORF532	ORF533	ORF534	ORF535	ORF536	ORF537	ORF538	ORF539	ORF540	ORF541	ORF542	ORF543	ORF544	ORF545

	begin	stop	Homology	Đ.	Species	Score	ě,
ORF546	533849	533244	YtqB	AF008220	Bacillus subtilis	273	88
ORF547	534672	533944	hypothetical protein MTCY08D5.03c	292669	Mycobacterium tuberculosis	170	35
ORF548	535915	534878	ribonucleoside diphosphate reductase, beta subunit (nrdB)	AE000553	Helicobacter pylori	397	33
ORF549	539153	535956	ribonucleoside-diphosphate reductase l alpha subunit (nrdA)	AE000581	Helicobacter pylori	1447	51
ORF550	539731	540519	phosphatidylserine synthase (pssA)	AE000614	Helicobacter pylori	226	49
ORF551	540523	540969	putative				
ORF552	540906	541805	hypothetical 54.7 kD protein in udp 3' region precursor (0475)	AE000459	Escherichia coli	82	33
ORF553	543255	541825	Ydr430cp; CAI: 0.15	U33007	Saccharomyces cerevisiae	130	48
ORF554	544133	543222	putative				
ORF555	544565	544179	hypA gene product	X86493	Clostridium perfringens	221	46
ORF556	544762	544487	orfl gene product	X70951	Saccharomyces cerevisiae	153	æ
ORF557	546423	544951	serine protease (htrA)	AE000610	Helicobacter pylori	981	46
ORF558	547480	546584	succinyl coenzyme A synthetase alpha subunit	U23408	Dictyostelium discoideum	869	63
ORF559	546789	547382	putative				T
ORF560	547901	547476	putative succinyl-coA synthetase beta chain	AJ000975	Bacillus subtilis	388	55
ORF561	548634	547900	succinateCoA ligase (ADP-forming)	X54073	Thermus aquaticus flavus	498	46
ORF562	548692	549459	cell division protein (ftsY)	AE000588	Helicobacter pylori	330	46
ORF563	550385	549663	putative				

18	40	36	45	51					39	38	45	!	39	45	42			35		83
Score	508	353	1324	1009					245	130	519		874	594	334			203		315
Species	Escherichia coli	Haemophilus influenzae	Thermus aquaticus thermophilus	Haemophilus influenzae					Chlamydia psittaci	ı	Bacillus subtilis		Escherichia coli	Cucumis sativus	Drosophila melanogaster			Mycobacterium tuberculosis		Chlamydia trachomatis
ar	D90832	U32730	U17352	U32824					U65942	U72499	Y13937		U18997	M80571	M58465			294752		M62820
Homology	Tyrosine-specific transport protein (Tyrosine permease).	tyrosine-specific transport protein (tyrP)	L-glutamine:D-fructose-6-P amidotransferase precursor	hypothetical	putative	putative	putative	putative	POMP91A	putative 98 kDa outer membrane protein	putative PlsX protein	1	ORF_f495; orfF of ECMRED, uses 2nd start	glycerol-3-phosphate acyltransferase	insulin-degrading enzyme	putative	putative	unknown	putative	putative heat shock protein ORF; putative
stop	550421	551797	553096	554927	556904	557314	558235	558310	961655	561150	121899	563943	566953	567966	570399	572021	572755	572731	573427	573660
pegin	551611	553041	554946	556300	556524	558126	557810	559215	561349	562931	564083	563593	565379	567079	568021	571269	572519	573519	572879	574160
ORF	ORF564	ORF565	ORF566	ORF567	ORF568	ORF569	ORF570	ORF571	ORF572	ORF573	ORF574	ORF575	ORF576	ORF577	ORF578	ORF579	ORF580	ORF581	ORF582	ORF583

<b>*</b>	66	39	78	81	37	48	48	66	T				41	41		42	25	46	42
Score	384	176	358	393	94	695	243	5054					298	339		673	845	719	156
Species	Chlamydia trachomatis	Helicobacter pylori	Chlamydia trachomatis	Chlamydia trachomatis	Bacillus subtilis	Synechocystis sp.	Bacillus subtilis	Chlamydia trachomatis					Staphylococcus aureus	Listeria monocytogenes		Synechocystis sp.	Aquifex pyrophilus	Haemophilus influenzae	Rhodobacter sphaeroides
a	M62820	AE000630	U31570	U31570	X16518	D90899	U87792	U20547					AB001896	U13165		D90909	U71154	U32691	U29587
Homology	ribosomal protein S18		peptidyl-tRNA hydrolase	peptidyl-tRNA hydrolase	partial ctc gene product (AA 1-186)	glycogen (starch) synthase	phosphatidylglycerophosphate synthase	glycyl-tRNA synthetase	putative	putative	putative	putative	dnaG	DNA primase	putative	DNA mismatch repair protein	DNA mismatch repair protein	excinuclease ABC subunit C (uvrC)	exinuclease ABC subunit C
stop	574184	574446	574923	575057	575469	578023	578017	582104	582206	582811	583182	583438	583827	584299	585016	586610	587758	589408	589578
begin	574426	574781	575243	575458	575849	576545	578673	579012	582697	583122	583514	583869	584435	584967	585297	585240	586484	587786	589198
ORF	ORF584	ORF585	ORF586	ORF587	ORF588	ORF589	ORF590	ORF591	ORF592	ORF593	ORF594	ORF595	ORF596	ORF597	ORF598	ORF599	ORF600	ORF601	ORF602

1%			8	T	Τ	49	53	52		38	42	42			Τ		45	49	37	25	g	Τ	41
	-	-	$\perp$	╁	-		+	-	+	+-		+			+	+	4	4	F.	<del> </del>	\frac{\cap_{\cap_{\chi}}}{\chi_{\chi}}	$\perp$	4
Score			117			991	375	571		1097	242	388			:		254	638	291	1069	196		780
Species			Bacillus subtilis			Haemophilus influenzae	Synechocystis sp.	Thermus aquaticus thermophilus		Bacillus subtilis	Bacillus subtilis	Escherichia coli					Oryza sativa	Thermoactinomyces intermedius	Arabidopsis thaliana	Ricinus communis	Escherichia coli		Helicobacter pylori
ΩĮ			X62539			U32693	D90906	X70708		X13937	U51868	L14681					U33283	X79068	X57545	L13242	U82598		AE000579
Homology	putative	putative	homologous to E.coli rnpA	putative	putative	cys-tRNA synthetase (cysS)	lysyl-tRNA synthetase	lysinetRNA ligase	putative	putative PriA protein	L-alanine - pimelyl CoA ligase	2-	acylglycerophosphoethanolami ne acyltransferase/acyl	carrier protein synthetase	putative	putative	3'(2'),5-diphosphonucleoside 3'(2') phosphohydrolase	leucine dehydrogenase	inorganic pyrophosphatase	beta-ketoacyl-ACP synthase	HI0034 homolog	putative	conserved hypothetical protein
stop	589630	591272	592765	592849	593121	595637	595640	596154	597282	608009	600734	601910			603531	603757	605610	605582	607493	608031	609296	610109	612927
begin	590061	590739	592406	593145	593900	594138	596122	. 596864	597731	598524	601876	603523			603794	604413	604549	606619	606843	890609	609652	611860	611812
ORF	ORF603	ORF604	ORF605	ORF606	ORF607	ORF608	ORF609	ORF610	ORF611	ORF612	ORF613	ORF614			ORF615	ORF616	ORF617	ORF618	ORF619	ORF620	ORF621	ORF622	ORF623

*	£.	ස 		31				55	47		46	47	47	56	39	46			40	43	42
Score		243		136				467	278		361	460	756	436	1121	1426			416	638	283
Species		Synechocystis sp.		Thermoanaerobacter				Escherichia coli		- 1	Bacillus subtilis	Bacillus subtilis	Haemophilus influenzae	Rhizobium leguminosarum	Thiobacillus ferrooxidans	Haemophilus influenzae			Bacillus subtilis	Salmonella typhimurium	Synechocystis sp.
ΩĪ	L	X62437		U56021				U14003	D90837		273234	Z73234	U32783	X59956	X95571	U32805			M97208	U06779	D90912
Homology	Imp_N_coct;:	glutamyl-2, 6-	diaminopimelateD-alanyl-D- alanine ligase	chaperonin 60	putative	putative	putative	elongation factor P	AMP nucleosidase (EC	1.2.2.1.	transketolase	transketolase	transketolase 1 (TK 1) (tktA)	alanyl-tRNA synthetase	alanyl-tRNA synthetase	transcription-repair coupling factor (trcR) (mfd)	1	putative	uroporphyrinogen decarboxylase	putative oxygen-independent coproporphyrinogen III oxidase	oxygen independent coprophorphyrinogen III oxidase
stop	624073			626665	626900	627853	628124	628146	629801	100003	629804	630298	630915	638084	640207	643472	0.00	640220	644495	645430	645840
hegin	624918			625346	626514	626954	627822	628715	628932	630406	930400	630960	631799	637488	638036	640221	20000	179049	643485	644471	645394
ORF	ORF638			ORF639	ORF640	ORF641	ORF642	ORF643	ORF644	OPF645	OF OF O	ORF646	ORF647	ORF648	ORF649	ORF650	100000	OKFOST	ORF652	ORF653	ORF654

000	Poort						
33000	Degin	dons	HomoLogy	Ħ	Species	Score	%I
OKF655	645840	647111	hemY	M97208	Bacillus subtilis	133	88
OKF656	649676	647109	phosphoprotein	L25078	Chlamydia trachomatis	2043	66
ORF657	649970	650344	нст	M60902	Chlamydia trachomatis	603	100
ORF658	650418	651722	pCTHoml gene product	M94254	Chlamydia trachomatis	1735	100
ORF659	651686	652171	putative				
ORF660	652516	652908	phenolhydroxylase component	U32702	Haemophilus influenzae	263	41
ORF661	652799	653593	phenolhydroxylase component	U32702	Haemophilus influenzae	456	51
ORF662	659884	661851	YtpT	AF008220	Bacillus subtilis	709	52
ORF663	661740	662282	spollIEB protein	M17445	Bacillus subtilis	330	43
ORF664	662286	663074	уусд	D78193		405	38
ORF665	662951	663730	C41G7.4	281048	Caenorhabditis elegans	200	36
ORF666	664212	663745	hypothetical protein MTCY180.08	297193	Mycobacterium tuberculosis	194	38
ORF667	665619	664255	D-alanine glycine permease (dagA)	AE000603	Helicobacter pylori	205	¥
ORF668	680999	665727	putative				
ORF669	666423	665782	putative				
ORF670	666831	668117	putative				
ORF671	668121	668375	putative				
ORF672	668470	668174	riboflavin synthase beta chain (ribE)	U32810	Haemophilus influenzae	192	40
ORF673	669533	668616	GTP cyclohydrolase II / 3,4-dihydroxy-2-butanone-4-phoshate synthase	AJ000053	Arabidopsis thaliana	800	51
ORF674	669892	669485	unnamed protein product	A38767	Saccharomyces	288	64
ORF675	670780	866699	ribG gene product	L09228	Bacillus subtilis	191	42
						-	•

671241         670732         riboflavin-specific         U27202           671182         672447         seryl-tRNA synthetase         X91007           672692         673231         putative         128104           673204         674562         ATPase         128104           673204         674562         ATPase         128104           673204         674562         ATPase         128104           673204         674562         ATPase         128104           673204         675327         unknown         274025           675327         676463         rod-shape-determining         M22857           675327         676476         biotin lacetyl-CoA         L02354           677027         676476         biotin lacetyl-CoA         L02354           67842         677027         676476         biotin lacetyl-CoA         L02354           67842         67871         680502         syntative         AE000542           68153         680559         putative         M2718           68154         681465         ORF 4         M2718           68164         684418         putative         Bettide, 505 asl           6817         681936         <		(	ě
672447 seryl-tRNA synthetase 672447 seryl-tRNA synthetase 673231 putative 674562 ATPase 675232 unknown 676463 rod-shape-determining protein 676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 679508 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster 681280 Nifu 682558 putative 684165 ORF 4 684165 ORF 4 684165 ORF 4 684109 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688193 putative 688132 putative 688132 putative 688132 putative 688132 putative 689146 putative	1	Score	<b>₽</b>
672447 seryl-tRNA synthetase 673231 putative 674562 ATPase 675232 unknown 676463 rod-shape-determining 676476 biotin [acetyl-CoA 677700 ORFX13 679508 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifs) 681280 NifU 682558 putative 683087 putative 684465 ORF 4 684465 ORF 4 684465 ORP 4 684418 putative 684203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688193 putative 688193 putative 688132 putative 688132 putative 688133 putative 688132 putative 688131 putative	2 Actinobacillus   pleuropneumoniae	314	51
673231 putative 674562 ATPase 675232 unknown 676463 rod-shape-determining 676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 679508 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifs) 681280 NifU 682558 putative 683087 putative 684465 ORF 4 684465 ORF 4 684465 ORF 4 68418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688150 putative 688153 putative 688153 putative 688169 putative 689846 putative		736	49
675232 unknown 676463 rod-shape-determining protein 676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 680502 synthesis of [Fe-S] cluster (B8258 putative 683087 putative 684465 ORF 4 684418 putative 686203 ayrchesis of [human, infertile patient, testis, Peptide, 505 aa] 687204 b-glycerol 3-phosphate 688150 putative 688150 putative 688150 putative 688150 putative 688150 putative 688160 putative 688160 putative 688160 putative 688160 putative 688160 putative		_	
675232 unknown 676463 rod-shape-determining protein 676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 689502 synthesis of [Fe-S] cluster (nifs) 681280 NiffU 682558 putative 683087 putative 684465 ORF 4 684465 ORF 4 684465 ORF 4 686203 Agx-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688193 putative 688193 putative 688432 putative 688432 putative 689631 putative 689643 putative	4 Transposon Tn5422	565	41
676463 rod-shape-determining protein 676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 680502 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifS) 681280 NifU 682558 putative 682465 ORF 4 684465 ORF 4 684465 ORF 4 684465 ORF 4 68418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688360 putative 688193 putative 688432 putative 688432 putative 689846 putative		340	43
676476 biotin [acetyl-CoA carboxylase] ligase 677700 ORFX13 679508 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifs) 681280 NiffU 682558 putative 683087 putative 684465 ORF 4 684465 ORF 4 684465 ORF 4 684418 putative 688410 L-glycerol 3-phosphate 688150 putative 688193 putative 688132 putative 688132 putative 688132 putative 689631 putative 689643 putative		442	37
Carboxylase  ligase	1 Paracoccus	169	49
677700 ORFX13 680502 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifS) 681280 NifU 68258 putative 683087 putative 684465 ORF 4 684418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688150 putative 688132 putative 688432 putative 6886432 putative 689641 putative 689643 putative		}	!
679508 2,3-bisphosphoglycerate 680502 synthesis of [Fe-S] cluster (nifS) 681280 NifU 68258 putative 688465 ORF 4 684465 ORF 4 688418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688150 putative 688193 putative 688432 putative 6886432 putative 689631 putative 689646 putative	Bacillus subtilis	426	43
680502 synthesis of [Fe-S] cluster (nifS) 681280 NifU 682558 putative 683087 putative 684465 ORF 4 684418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate 688350 putative 688193 putative 688432 putative 689631 putative 689646 putative		494	47
682558 putative 683087 putative 684465 ORF 4 684418 putative 686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-Glycerol 3-phosphate 688150 putative 688193 putative 688432 putative 689631 putative 6896463 putative	342 Helicobacter pylori	150	33
684465 putative 684465 ORF 4 688418 putative 6886203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate dehydrogenase 688193 putative 688432 putative 689631 putative 689631 putative 689643 putative 689643 putative 690463 putative	780 Cyanothece PCC 8801	101	31
684465 ORF 4  684418 putative  686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa]  687204 L-glycerol 3-phosphate dehydrogenase 688360 putative 688193 putative 688432 putative 689631 putative 689631 putative 689631 putative 6896463 putative			
684465 ORF 4  684218 putative  686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa]  687204 L-glycerol 3-phosphate dehydrogenase 688360 putative 688432 putative 688432 putative 689631 putative 689643 putative			
684418         putative           686203         AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa]           687204         L-glycerol 3-phosphate dehydrogenase           688360         putative           688193         putative           688432         putative           689631         putative           689646         putative           689646         putative	Bacillus subtilis	70.8	36
686203 AgX-1 antigen [human, infertile patient, testis, Peptide, 505 aa] 687204 L-glycerol 3-phosphate dehydrogenase 688150 putative 688193 putative 688432 putative 689631 putative 6896463 putative 6896463 putative		3	3
687204 L-glycerol 3-phosphate 688360 putative 688193 putative 688432 putative 689631 putative 6896463 putative	Homo sapiens	338	37
687204       L-glycerol 3-phosphate         dehydrogenase         688360       putative         688193       putative         688432       putative         689631       putative         689846       putative         68963       putative			
688193 688193 688432 689631 689846 690463	Escherichia coli	577	88
688193 688193 688432 689631 689846 690463			-
688193 688432 689631 689846 690463			
688432 689631 689846 690463			
689631 689846 690463			
689846			
690463			
691871 690672 putative			

ORF	begin	Btop	Homology	ET.	Contracto		Ş
005500	602037	10000	(6)-10-10-10-10-10-10-10-10-10-10-10-10-10-		Sarcade	Score	<b>9</b>
OOLING	150560	032041	pnospnoenolpyruvate carboxykinase	M59372	Neocallimastix frontalis	1818	29
ORF701	694934	693837	MreB protein	M96343	Bacillus subtilis	961	56
ORF702	697263	694942	SNF	X98455	Bacillus cereus	1073	20
ORF703	698084	697170	putative		1		
ORF704	698392	626269	putative				
ORF705	698792	700117	trigger factor (tig)	AE000591	Helicobacter pylori	84	34
ORF706	700269	700895	proteosome major subunit	AF013216	Myxococcus xanthus	615	59
ORF707	700912	702165	ATP-dependent protease	L18867	Escherichia coli	1183	55
ORF708	702183	703412	poly(A) polymerase	L47709	Bacillus subtilis	362	38
ORF709	703522	705000	hypothetical protein	D90912	Synechocystis sp.	809	14
ORF710	705011	705604	putative				
ORF711	706159	705704	Preprotein translocase subunit	AF022186	Cyanidium caldarium	165	44
ORF712	706521	706138	secA	X99401	Bacillus firmus	155	42
ORF713	708103	706496	K 200	110003		CCT	7
		00.	V Sector	066081	Mycobacterium smegmatis	1044	 82
ORF714	708398	708078	cp-SecA; chloroplast SecA homolog	U71123	Zea mays	258	69
ORF715	708610	708248	SecA	U21192	Streptomyces lividans	179	42
ORF716	710278	708872	putative				
ORF717	711164	710262	phosphatidylserine decarboxylase	U72715	Chlamydia trachomatis	1548	66
ORF718	711432	712763	homologous to E.coli 50K	X62539	Bacillus subtilis	713	8
ORF719	712767	713438	ultraviolet N-glycosylase/AP	U22181	Micrococcus luteus	273	45
ORF720	714232	713651	Tyase				
ORF721	714632	714120	putative				
ORF722	715592	714834	putative				
ORF723	715854	715558	putative				
ORF724	716937	715921	putative				

47	┿		45	T	37	41	4	20	39	4	22	85	77	55		33	43	66	76	06	46
Score	2049		997		164	723	477	388	805	154	607	266	854	531		115	208	2045	1269	1278	1153
Species	Chlamydia trachomatis		Haemophilus influenzae		Haemophilus influenzae	Bacillus subtilis	1.0	Caulobacter crescentus	Escherichia coli	1.4	10	Synechocystis sp		,		Saccharomyces cerevisiae	Haemophilus influenzae	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis
A	U83197		U32834		U32834	X56678	U00039	U87804	D90811	D64004	D90811	D64004	D64004	D64004		L36940	U32688	M14738	U60196	U60196	U60196
Homology	3-phosphoglycerate kinase	putative	phosphate permease (YBR296C)	putative	H. influenzae predicted coding region H11603	dciAD	мая фррЕ	chromosome partitioning protein ParB	Nifs protein.	hypothetical protein	Multidrug resistance protein 1 (P-glycoprotein 1).	ABC transporter subunit	ABC transporter subunit	ABC transporter subunit	putative	antiviral protein	penicillin-binding protein 2 (pbp2)	major outer membrane protein precursor	ribosomal protein S2	elongation factor Ts	UMP kinase
stop	717149	718862	718499	719782	720144	721575	722356	722397	723378	724576	725767	726538	726753	727469	728329	728759	729442	734427	735659	736504	737254
begin	718357	718500	719797	720273	720452	720613	721559	723248	724598	725763	726519	726819	727493	727984	728778	729346	732639	733246	734814	735644	736520
ORF	ORF725	ORF726	ORF727	ORF728	ORF729	ORF730	ORF731	ORF732	ORF733	ORF734	ORF735	ORF736	ORF737	ORF738	ORF739	ORF740	ORF741	ORF742	ORF743	ORF744	ORF745

48	35		31	37			40	38					55							40	38	49	8
Score	760		116	453			137	117					1220							172	970	409	107
Species	Chlamydia trachomatis		Listeria monocytogenes	Pseudomonas aeruginosa			Mycobacterium tuberculosis	Caulobacter crescentus					Pseudomonas aeruginosa							Bacillus subtilis	Clostridium acetobutylicum	Mycobacterium smegmatis	Mycobacterium tuberculosis
Ð	U60196		U40604	X68594			280233	M69228					AF010151							M57676	U35453	X92503	274024
Homology	ribosome-releasing factor	putative	ORF3; putative 39 kDa protein	XcpQ	putative	putative	unknown	putative	putative	putative	putative	putative	PscN	putative	putative	putative	putative	putative	putative	NAD(P)H:glutamyl-transfer RNA reductase	DNA gyrase subunit B	gyrA	unknown
stop	737787	738679	739740	740060	742045	742824	743306	744430	744611	744958	745561	746416	746944	748274	748965	749433	749721	750007	752503	753616	756814	758301	758446
begin	737254	737942	738838	742057	742869	743378	744298	744714	744985	745557	746412	746772	748269	748966	749426	749702	750029	752307	752913	754659	755000	756796	758691
ORF	ORF746	ORF747	ORF748	ORF749	ORF750	ORF751	ORF752	ORF753	ORF754	ORF755	ORF756	ORF757	ORF758	ORF759	ORF760	ORF761	ORF762	ORF763	ORF764	ORF765	ORF766	ORF767	ORF768

SfhB         U50134         Escherichia coli           putative         042493         Chlamydia trachomatis           8-phosphate synthetase         U72493         Chlamydia trachomatis           anknown         U72493         Chlamydia trachomatis           ATP binding protein         U72493         Chlamydia trachomatis           chlanectin coding region         M17875         Chlamydia trachomatis           unknown function         Z32530         Chlamydia trachomatis           putative         Dutative         Dutative           putative         Dutative         Bacillus subtilis           putative         U18997         Escherichia coli           DNA topolsomerase I         L27797         Bacillus subtilis           putative         Dubative         Dubative	ORF	pegin	stop	Homology	ŧ	20,0000	1	
760242         75991         putative           760548         Putative         760588         Putative           760568         761772         3-deoxy-D-manno-octulosonate (72493)         Chlamydia trachomatis           761759         762142         unknown         U72493         Chlamydia trachomatis           762267         762083         ATP binding protein         U72493         Chlamydia trachomatis           764657         76438         putative         M17875         Chlamydia trachomatis           76668         764857         764857         Chlamydia trachomatis           76668         764857         Chlamydia trachomatis           76668         768091         RecA         U16739         Chlamydia trachomatis           766681         766055         unknown function         Z32530         Chlamydia trachomatis           770032         770033         770034         Putative         T70039           770034         770034 </td <td>F769</td> <td>759787</td> <td>759338</td> <td></td> <td>US0134</td> <td></td> <td>Score</td> <td>27</td>	F769	759787	759338		US0134		Score	27
76058   760188   putative   76058   76172   3-deoxy-D-manno-octulosonate   172493   Chlamydia trachomatis     761759   762142   unknown   172493   Chlamydia trachomatis     762267   762983   ATP binding protein   172493   Chlamydia trachomatis     764857   764438   putative   176643   766068   76421   unknown function   232530   Chlamydia trachomatis     766643   76605   unknown function   232530   Chlamydia trachomatis     76608   768791   unknown function   232530   Chlamydia trachomatis     770092   766791   unknown function   232530   Chlamydia trachomatis     770040   770438   putative   232530   Chlamydia trachomatis     770051   770430   putative   232530   Chlamydia trachomatis     770052   770430   putative   232530   Chlamydia trachomatis     770053   770430   putative   232530   Chlamydia trachomatis     77054   770534   putative   232530   Chlamydia trachomatis     77054   770450   putative   232530   Chlamydia trachomatis     77054   770450   putative   232530   Chlamydia trachomatis     77054   770450   putative   232530   Chlamydia trachomatis     77054   770534   putative   232530   Chlamydia trachomatis     770553   770553   putative   232530   Chlamydia trachomatis     770554   7705551   Putative   232530   Chlamydia trachomatis     7705551   770553   Putative   232530   Chlamydia trachomatis     770551   770553   Putative   232530   Chlamydia trachomatis     770551   770553   Putative   232530   Putative   232530   Putative   232530   Putative   232530   Putative   2325	F770	760242	759871	putative			147	10
760966   761772   3-deoxy-D-manno-octulosonate   U72493   Chlamydia trachomatis     761759   762142   unknown   U72493   Chlamydia trachomatis     762267   762983   ATP binding protein   U72493   Chlamydia trachomatis     764857   764438   Putative   U16068   764821   unknown function   Z32530   Chlamydia trachomatis     766068   764821   unknown function   Z32530   Chlamydia trachomatis     766068   766934   RecA   U16739   Chlamydia trachomatis     76608   768791   V66065   unknown function   Z32530   Chlamydia trachomatis     768091   766954   RecA   U16739   Chlamydia trachomatis     770092   768791   unknown function   Z32530   Chlamydia trachomatis     7700128   770410   putative   Z32530   Chlamydia trachomatis     77024   770654   putative   D26185   Bacillus subtilis     77027   771391   unknown function   D26185   Bacillus subtilis     77037   771391   Unknown function   D26185   Bacillus subtilis     77663   77421   DNA topoisomerase   L27797   Bacillus subtilis     77663   77706   Putative   L27797   Bacillus subtilis     77653   77705   Putative   L27797   Bacillus subtilis     77705   Putative   L27797   Bacillus coli     77705   Putative   L27797   Putative   L27797   Bacillus coli     77705   Putative   L27797   Putative   L27797   Putative   L27797     7770	F771	760538	760188	putative				
761759 762142   unknown   U72493   Chlamydia trachomatis     762267 762983   ATP binding protein   U72493   Chlamydia trachomatis     764865 76335   Chlanectin coding region   M17875   Chlamydia trachomatis     764867 764438   putative   D16739   Chlamydia trachomatis     766068 764821   unknown function   Z32530   Chlamydia trachomatis     766043 766055   unknown function   Z32530   Chlamydia trachomatis     768091 766934   Rech   U16739   Chlamydia trachomatis     768092 768791   unknown function   Z32530   Chlamydia trachomatis     770092 768791   unknown function   Z32530   Chlamydia trachomatis     770138 770470   putative   D26185   Bacillus subtilis     770210 771330   Unknown function   D26185   Bacillus subtilis     772210 7713427   ORP_£169   U18997   Eacherichia coli     776035 774191   DAR topolsomerase I   L27797   Bacillus subtilis     776036 777195   D4Lative   D26185   Bacillus subtilis     776037 777195   776953   putative   D27797   Bacillus subtilis     777012 777195   776953   putative   D2818   Escherichia coli     777012 777195   777195   D4PLative   D7797   D4PLative     777195 777195   777195   D4PLative   D4PLative     777195 777195   777195   D4PLative   D4PLative     777195 777195   D4PLative   D4PLative   D4PLative     777195 777195   777195   D4PLative   D4PLativ	1F772	760966	761772	3-deoxy-D-manno-octulosonate 8-phosphate synthetase	i		1350	66
762267         762983         AIP binding protein         U72493         Chlamydia trachomatis           764465         763335         chlanectin coding region         M17875         Chlamydia trachomatis           764857         764821         unknown function         232530         Chlamydia trachomatis           766068         764821         unknown function         232530         Chlamydia trachomatis           766043         766934         RecA         U16739         Chlamydia trachomatis           768091         766934         RecA         U16739         Chlamydia trachomatis           768092         768785         unknown function         232530         Chlamydia trachomatis           770032         768785         unknown function         232530         Chlamydia trachomatis           770032         770418         putative         232530         Chlamydia trachomatis           770641         770641         770641         putative         770641           77018         770641         770641         putative         776618           77603         77703         putative         1627797         Bacillus subtilis           77603         77703         putative         1627797         Bacillus subtilis <t< td=""><td>E773</td><td>761759</td><td>762142</td><td>unknown</td><td>U72493</td><td></td><td>536</td><td>94</td></t<>	E773	761759	762142	unknown	U72493		536	94
764465         763335         chlanectin coding region         M17875         Chlamydia trachomatis           764857         764438         putative         232530         Chlamydia trachomatis           766068         764821         unknown function         232530         Chlamydia trachomatis           766091         766043         RecA         U16739         Chlamydia trachomatis           768091         768785         768781         unknown function         232530         Chlamydia trachomatis           770092         768791         unknown function         232530         Chlamydia trachomatis           770018         770018         putative         232530         Chlamydia trachomatis           770018         77018         putative         232530         Chlamydia trachomatis           770018         77018         putative         232530         Chlamydia trachomatis           770018         77018         putative         232530         Chlamydia trachomatis           77001         77018         putative         232530         Chlamydia trachomatis           77021         77018         putative         232530         Chlamydia trachomatis           77021         77023         774230         774230         7742	F774	762267	762983		U72493	l l	1197	66
764857         764438         putative         232530         Chlamydia trachomatis           766068         764821         unknown function         232530         Chlamydia trachomatis           766043         766043         RecA         U16739         Chlamydia trachomatis           768091         76804         RecA         U16739         Chlamydia trachomatis           768092         768791         unknown function         Z32530         Chlamydia trachomatis           770092         768791         unknown function         Z32530         Chlamydia trachomatis           770138         770138         770470         putative         Chlamydia trachomatis           77054         77054         77054         Putative         D26185         Bacillus subtilis           772010         771330         putative         U18997         Bscherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776653         777603         putative         Bacillus subtilis           777195         777552         777732         ORP_1397         U29581         Bscherichia coli	F775	764465	763335	chlanectin coding region	M17875	Chlamydia trachomatis	239	100
766068         764821         unknown function         Z32530         Chlamydia trachomatis           766643         766065         unknown function         Z32530         Chlamydia trachomatis           768091         768052         unknown function         Z32530         Chlamydia trachomatis           768185         768791         unknown function         Z32530         Chlamydia trachomatis           770092         768791         unknown function         Z32530         Chlamydia trachomatis           770018         770138         770418         putative         Chlamydia trachomatis           770541         770541         putative         D26185         Bacillus subtilis           772300         773391         unknown         U18997         Escherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           77663         7771663         777195         putative         D173797         Bacillus subtilis           777195         777195         777195         Packerichia coli         C77197         Bacillus subtilis           777195         777195         777797         Bacillus subtilis         C77197         Bacillus coli           777195         777195	F776	764857	764438	putative				
766643         766065         unknown function         Z32530         Chlamydia trachomatis           768091         768092         RecA         U16739         Chlamydia trachomatis           768185         768252         unknown function         Z32530         Chlamydia trachomatis           770092         768791         unknown function         Z32530         Chlamydia trachomatis           770138         770470         putative         Chlamydia trachomatis           770561         770185         putative         D26185           772010         771330         unknown         D26185         Bacillus subtilis           772390         774221         773427         ORE_f169         U18997         Escherichia coli           776635         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776636         777105         Putative         CALARA COLI           777210         777195         Putative         CALARA COLI           777195         777195         Putative         CALARA COLI           777195         777195         Putative         CALARA COLI           777195         777197         CALARA COLI           7771195         7777197         CALARA	E777	766068	764821	unknown function	232530	Chlamydia trachomatis	1803	66
768091         766934         RecA         U16739         Chlamydia trachomatis           768785         768792         unknown function         Z32530         Chlamydia trachomatis           770092         768791         unknown function         Z32530         Chlamydia trachomatis           770138         770470         putative         Chlamydia trachomatis           770661         770185         putative         Chlamydia trachomatis           770924         770185         putative         Chlamydia trachomatis           772010         771330         putative         Chlamydia trachomatis           772390         773391         unknown         D26185         Bacillus subtilis           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           77663         777106         putative         Bacillus subtilis           77663         777106         putative         Bacillus subtilis           777122         777123         ORF_f1997         Bccherichia coli           777122         777125         Robertichia coli	E778	766643	766065	unknown function	Z32530	1	704	100
768785         768252         unknown function         Z32530         Chlamydia trachomatis           770092         768791         unknown function         Z32530         Chlamydia trachomatis           770138         770470         putative         Chlamydia trachomatis           770924         770661         770634         putative           772300         771330         putative         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Bacillus subtilis           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776633         777106         putative         Bacillus subtilis           776633         777106         putative         Bacillus subtilis           777105         776036         Recherichia coli	F779	768091	766934	RecA	U16739	1	1753	100
770092         768791         unknown function         Z32530         Chlamydia trachomatis           770138         770470         putative         C70661         77084         putative           770924         770534         putative         D26185         Bacillus subtilis           772390         773391         unknown         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Escherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           77663         777105         putative         Bacillus subtilis           777195         77653         putative         CATTATATATATATATATATATATATATATATATATATA	F780	768785	768252	unknown function	Z32530		904	66
770138         770470         putative           770661         770185         putative           770924         770134         putative           772390         773391         unknown         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Escherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776663         777106         putative         Bacillus subtilis           777195         776953         putative         Escherichia coli           777122         777732         ORF_f397         U29581         Escherichia coli	F781	770092	768791	unknown function	Z32530		2249	100
770661         770185         putative           770924         770634         putative           772390         773391         unknown         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Escherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           77663         777106         putative         Bacillus subtilis           777195         776953         putative         Bacillus subtilis           777122         777132         ORF_f397         U29581         Escherichia coli	F782	770138	770470	putative				
770924         770634         putative           772010         771330         putative           772390         773391         unknown         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Bscherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776663         777106         putative         Bacillus subtilis           777105         776953         putative         Bacillus subtilis           777105         777105         Putative         Bacillus subtilis	F783	770661	770185	putative				
772010         771330         putative         D26185         Bacillus subtilis           772390         773391         unknown         D26185         Bacillus subtilis           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776663         777105         putative         Acherichia coli           777195         777132         ORF_f397         U29581         Escherichia coli	F784	770924	770634	putative				
772390         773391         unknown         D26185         Bacillus subtilis           774221         773427         ORF_f169         U18997         Bscherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776663         777105         putative         Pacillus subtilis           777195         776953         putative         Bacillus subtilis           777195         777132         ORF_f397         U29581         Escherichia coli	785	772010	771330	putative				
774221         773427         ORF_f169         U18997         Escherichia coli           776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           776663         777706         putative         Putative           777195         777732         ORF_f397         U29581         Escherichia coli	5786	772390	773391	unknown	D26185		486	35
776035         774191         DNA topoisomerase I         L27797         Bacillus subtilis           77663         777106         putative         A77195         776953         putative           779222         777732         ORF_f397         U29581         Escherichia coli	F787	774221	773427	ORF_f169	U18997	Escherichia coli	263	51
777195 777732 putative 779222 777732 ORF_f397 U29581 Escherichia coli	F788	776035	774191	isomerase	L27797	Bacillus subtilis	1357	52
777195 776953 putative 779222 777732 ORF_f397 U29581 Escherichia coli	F789	776663	777706	putative				
779222 777732 ORF_f397 U29581 Escherichia coli	F790	777195	776953	putative				
	F791	779222	777732	ORF_£397	U29581	Escherichia coli	93	40

4			49		8	42	!		33	T	37		52				32	36	88	37		51	46			
Score			557		81	700	)		84		409		867				103	216	184	107		290	150		•	
Species			Escherichia coli		Escherichia coli	Gallus gallus			Rickettsia prowazekii		Vibrio alginolyticus	•	Bradyrhizobium japonicum				Neisseria meningitidis	Methanococcus jannaschii	Bacillus subtilis	Bacillus Stearothermophilus		Escherichia coli	Mycoplasma-like	organism		
ΩÏ			X04581		M31792	M12105			U02878		237111		124386				X92405	U67553	D84432	D13293		X61000	L22217			
Homology	putative	putative	exonuclease V (AA 1-1180)	putative	MreC protein	aspartate aminotransferase	precursor	putative	GreA	putative	NADH:ubiquinone	oxidoreductase subunit A	delta_aminolevulinic acid dehydratase	putative	putative	putative	ompR gene product	glucose-1-phosphate thymidylyltransferase	YqiD	farnesyl diphosphate synthase	putative	Orf39.9	This ORF is homologous to a	40.0 kd hypothetical protein	in the htrB 3' region from E. coli. Accession Number	X61000
stop	781552	782442	785524	786002	785546	786611		788021	787920	790609	792016		792059	794056	794957	795144	796255	797015	797365	797856	798086	797935	798416			
pegin	779321	781297	782447	785532	786580	787741		787620	790124	790160	790634		793084	793343	794046	795401	795575	796278	796979	797260	797772	798426	798925			
ORF	ORF792	ORF793	ORF794	ORF795	ORF796	ORF797		ORF798	ORF799	ORF800	ORF801		ORF802	ORF803	ORF804	ORF805	ORF806	ORF807	ORF808	ORF809	ORF810	ORF811	ORF812			

%I	46	35	97	97	100	29		જ											38	32	100		45	!	48
Score	407	397	1716	973	280	775		125											150	90	2868		310		100
Species	Helicobacter pylori	Caenorhabditis elegans	Chlamydía trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia pneumoniae		Chlamydia pneumoniae											Odontella sinensis	Homo sapiens	Chlamydia trachomatis		Pisum sativum		Neisseria meningitidis
ΩI	AE000633	U40707	U50732	US0732	U50732	L23921		L23921											267753	AB002334	J05546		Y08611		X68068
Homology	ribosomal protein S4 (rps4)	apurinic/apyrimidinic endonuclease	mviB homolog	mviB homolog	lorf2; possible membrane- bound protein	76 kDa protein	putative	76 kDa protein	putative	30S ribosomal protein S20	KIAA0336	RNA polymerase sigma-subunit	putative	dihydropterin	pyrophosphokinase /dihydropteroate synthase	dihydropteroate synthase									
stop	799927	800029	802129	802673	803246	804220	805356	806282	808081	809008	620608	810328	812342	813522	813772	814334	814314	814396	815428	817456	819320	819713	820402		821061
begin	799301	800892	801062	802023	802851	803105	804307	805290	806453	808026	810461	811605	811725	812329	813455	813732	815213	814878	815733	816116	817608	819324	819704		820375
ORF	ORF813	ORF814	ORF815	ORF816	ORF817	ORF818	ORF819	ORF820	ORF821	ORF822	ORF823	ORF824	ORF825	ORF826	ORF827	ORF828	ORF829	ORF830	ORF831	ORF832	ORF833	ORF834	ORF835		ORF836

M58480 Thiobacil ferrooxio M63176 Staphyloc M63176 Staphyloc X73124 Bacillus U32712 Haemophil	M58480 M63176 M63176 X73124 X73124 L19954								
		M58480 M63176 M63176 X73124 U32712 L19954	M58480 M63176 M63176 X73124 X73124 U32712 L19954 U55047	M58480 M63176 M63176 X73124 X73124 U32712 U55047	M58480 M63176 M63176 X73124 U32712 U32712 L19954 Z34917	458480 463176 463176 173124 132712 13954 134917	M58480 M63176 M63176 M63176 U32712 U32712 L19954 L19954	M58480 M63176 M63176 X73124 X73124 U32712 U35047 Z34917	M58480 M63176 M63176 X73124 X73124 U32712 U35047
	f 1								
10 1 1 10.1 1 1	product e stretch (32 l to N-termir nal sequence tly cloned B.	abol	gene product ical sidue stretch (32 tical to N-termir signal sequence partly cloned B. gene.; putative ck protein GroEL	gene product ical sidue stretch (32 tical to N-termir signal sequence partly cloned B. gene.; putative ck protein GroEL	gene product  ical  cical  cical  cical to N-termir  signal sequence  partly cloned B.  gene.; putative  ck protein GroEL	gene product  ical  sidue stretch (32- tical to N-terminal signal sequence of partly cloned B. gene.; putative  ck protein GroEL  tein	gene product  ical  sidue stretch (32- tical to N-terminal signal sequence of partly cloned B. gene.; putative  ck protein GroEL  tein	gene product  ical  sidue stretch (32- tical to N-terminal signal sequence of partly cloned B. gene.; putative  ck protein GroEL  tein	gene product  ical  sidue stretch (32- tical to N-terminal signal sequence of partly cloned B. gene.; putative  ck protein GroEL  tein
regulator helicase helicase ipa-57d gene p putative hypothetical	regulator helicase helicase ipa-57d gene putative hypothetical 19/20 residue 51) identical putative signe unknown, partl	regulator helicase helicase ipa-57d gene putative hypothetical 19/20 residue 51) identical putative signe unknown, partl subtilis gene.	regulator helicase helicase ipa-57d gene putative hypothetical 19/20 residue 51) identical putative signe unknown, partl subtilis gene. heat shock pro	helicase helicase helicase putative hypothetical hypothetical putative signs unknown, partl subtilis gene. heat shock pro heat shock pro heat shock pro	regulator helicase helicase ipa-57d gene putative hypothetical hypothetical butative signs unknown, partl subtilis gene. heat shock pro heat shock pro putative putative putative putative putative	regulator helicase helicase ipa-57d gene putative hypothetical 19/20 residue 51) identical putative signs unknown, partl subtilis gene. heat shock pro heat shock pro putative putative putative putative putative	regulator helicase helicase ipa-57d gene putative hypothetical putative signs unknown, partl subtilis gene. heat shock pro heat shock pro putative	regulator helicase helicase helicase hypothetical hypothetical putative signs unknown, partl subtilis gene. heat shock pro heat shock pro putative	regulator helicase helicase ipa-57d gene putative hypothetical 19/20 residue 51) identical putative signs unknown, partl subtilis gene. heat shock pro heat shock pro putative
				826340 827014 827856 828007 829355 831119 832152 832744	826340 827014 827856 828007 829355 831119 832152 832152 833446	826340 827014 827856 828007 829355 831119 831119 8331446 833446 833802	826340 827014 827856 828007 829355 831119 832152 833744 833446 833802 833802	826340 827014 827856 828007 831119 832152 832152 833446 833446 8334679 8334679	
827250 putative 827230 hypothetical	827230 putative 827230 hypothetical 829275 19/20 residue 51) identical putative signe unknown, partl	827230 putative 827230 hypothetical 829275 19/20 residue 51) identical putative signe unknown, partl subtilis gene.	827230 hypothetical 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro	#27250 putative #27230 hypothetical #29275 19/20 residue 51) identical putative signs unknown, part1 #20953 heat shock pro #31748 bas1 protein #31751 putative	827250 putative 827230 hypothetical 829275 19/20 residue 51) identical putative signe unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832214 putative	#27250 putative #27230 hypothetical #29275 19/20 residue 51) identical putative signs unknown, partl subtilis general #31748 bas1 protein #31751 putative #32214 putative #33368 putative	827250 putative 827230 hypothetical 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 831748 bas1 protein 831751 putative 832214 putative 832805 putative 833869 putative	827250 putative 827230 hypothetical 829275 19/20 residue 51) identical putative signe unknown, part1 subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832214 putative 832805 putative 833879 putative 833879 putative	#27250 putative #27230 hypothetical #29275 19/20 residue 51) identical putative signé unknown, part1 subtilis gene. #330953 heat shock pro #31751 putative #33214 putative #33368 putative #33368 putative #33369 putative #334661 putative #335371 putative
82/230 nypotnetical	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene.	829275 19/20 residue 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro	829275 19/20 residue 829275 19/20 residue 51) identical putative signe unknown, partl subtilis gene. 830953 heat shock pro	829275 19/20 residue 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative	829275 19/20 residue 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative	829275 19/20 residue 829275 19/20 residue 51) identical putative signe unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative 833809 putative	829275 19/20 residue 829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 831748 bas1 protein 831751 putative 832214 putative 832805 putative 833879 putative 833879 putative 833879 putative	#29275 19/20 residue   829275 19/20 residue   51) identical   putative signé unknown, part   subtilis gene.   830953 heat shock pro   831748 bas1 protein   831751 putative   832805 putative   833869 putative   833869 putative   833861 putative   833861 putative   833861 putative   833879 putative   833861 putative   833879 putative   833877 putative   8338771 putative   833877 putative
	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene.	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative 833808 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832214 putative 8332805 putative 833368 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative 833805 putative 833868 putative 833869 putative	829275 19/20 residue 51) identical putative signs unknown, partl subtilis gene. 830953 heat shock pro 831748 bas1 protein 831751 putative 832805 putative 833879 putative 833879 putative 8334661 putative

*1	-																						$\prod$			47		98			40
Score																										206		444	-		181
Species	1																									Haemophilus influenzae		Bacillus subtilis			Pseudomonas aeruginosa
a																										U32692		D84432			L13865
Homology	putative	DNA mismatch repair protein (muth)	putative	YqhT	putative	putative	fimbrial assembly protein																								
stop	838699	839575	840583	841713	842459	843068	843031	846196	843802	846217	848150	850230	851669	853700	854920	855437	856233	857451	859587	860640	860724	861580	862098	863571	863996	866248	866605	867732	060698	869357	871372
begin	837209	838760	839942	840445	841659	842523	843495	843239	844137	848043	850123	851645	853696	854836	855525	856240	857183	859439	859946	859642	861599	862053	863540	863930	864697	864923	866303	599998	018298	869094	869270
ORF	ORF858	ORF859	ORF860	ORF861	ORF862	ORF863	ORF864	ORF865	ORF866	ORF867	ORF868	ORF869	ORF870	ORF871	ORF872	ORF873	ORF874	ORF875	ORF876	ORF877	ORF878	ORF879	ORF880	ORF881	ORF882	ORF883	ORF884	ORF885	ORF886	ORF887	ORF888

ORF	pegin	stop	Homology	គ	Species	Score	%I
ORF889	871299	872582	xpsE gene product	X59079	Xanthomonas campestris	825	26
ORF890	872429	872860	secretion protein XcpR	Y09102	Acinetobacter calcoaceticus	213	48
ORF891	872773	873915	ORF_0398	U18997	Escherichia coli	271	33
ORF892	873812	873360	putative				
ORF893	874028	874438	putative				
ORF894	874778	875386	putative				
ORF895	875774	876382	putative				
ORF896	877872	877000	secretion system apparatus, SsaT	X99944	Salmonella typhimurium	174	क्र
ORF897	878172	877876	yscs	L25667	Yersinia pseudotuberculosis	172	4
ORF898	879098	878172	pathogenicity protein	M64094	Xanthomonas campestris	464	46
ORF899	878883	879161	putative				
ORF900	879842	879105	Расъ	U56077	Pseudomonas aeruginosa	141	\$
ORF901	880885	880052	putative				
ORF902	881863	88088	HrcJ	U56662	Erwinia amylovora	236	43
ORF903	882904	881948	ORF YOR196c	275104	Saccharomyces cerevisiae	685	44
ORF904	883794	882901	dihydrolipoamide dehydrogenase	L31844	Clostridium magnum	578	38
ORF905	884296	883661	Υgiv	D84432	Bacillus subtilis	437	44
ORF906	884996	884508	putative				
ORF907	888777	885166	helicase of the snf2/rad54 family	D90916	Synechocystis sp.	824	43
ORF908	890172	888940	sodium-coupled branched- chain amino acid carrier	D49784	Clostridium perfringens	230	32
ORF909	891164	890325	putative Fmu protein	Y13937	Bacillus subtilis	220	14
ORF910	891463	911168	putative				
ORF911	893278	896168	DD-carboxypeptidase	M85047	Bacillus subtilis	302	39

18			39				43		52	47			51	86		<u>6</u>	76		37	40	T	43		SS.		39	59
Score			155				1974		1117	989			1339	1196		607	380		150	181		197		145		309	302
Species			Synechocystis sp.				Helicobacter pylori		Escherichia coli	Streptococcus	equisimilis		Synechocystis sp.	Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia psittaci		Synechocystis sp.	Staphylococcus aureus		Helicobacter pylori	1	Sinorhizobium meliloti		Pseudomonas aeruginosa	Synechocystis sp.
ΩŦ			D90908				AE000646		M17102	217214			D90910	X66126	1 1000	L40369	L39892		D90914	Z48003		U94318		L13845		U67855	D90910
Homology	putative	putative	hypothetical protein	putative	putative	putative	DNA polymerase III alpha-	subunit (dnaE)	UhpC protein	histidinetRNA ligase		putative	aspartyl-tRNA synthetase	mip-like protein	Ilono	o od s	trxA	putative	hypothetical protein	DNA polymerase III	putative	VdlD	putative	acid-inducible gene	putative	UDP-3-0-acyl-GlcNAc deacetylase	(3R)-hydroxymyristol acyl carrier protein dehydrase
stop	893808	893643	893821	894248	895050	896829	897064		900791	903876		903471	905605	906474	906945	C#6006	100406	908742	909194	909584	909951	695016	910944	912261	912629	913218	913676
pegin	893356	893909	894276	894778	895892	895951	900783		902032	902659		903731	903860	905725	406493	50500	907306	908101	908721	861606	909583	180016	519016	910948	912399	912595	913203
ORF	ORF912	ORF913	ORF914	ORF915	ORF916	ORF917	ORF918		ORF919	ORF920		ORF921	ORF922	ORF923	ORF924	170	ORF925	ORF926	ORF927	ORF928	ORF929	ORF930	ORF931	ORF932	ORF933	ORF934	ORF935

e 18	+	42				48	43	╁	48	88	49	22	62	86	100	100	100	8
Score	503	407		-		470	210	116	800	315	240	605	434	343	419	618	568	100
Species	Rickettsia rickettsii	Escherichia coli				Haemophilus influenzae	Synechococcus sp.	IN	Mycoplasma-like organism	Mycoplasma pneumoniae	Thermotoga maritima	Haemophilus influenzae	Thermotoga maritima	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamadia transfer
Ð	L22690	x63666				U32761	AB000111	221677	M74770	AE000061	221677	U32761	221677	M80325	M80325	M80325	M80325	Menane
Homology	UDP-N-acetylglucosamine acyltransferase	methionyl-tRNA formyltransferase	putative	putative	putative	ribosomal protein L3 (rpL3)	50S ribosomal protein L4	ribosomal protein L23	rp12	Mycoplasma pneumoniae, ribosomal protein S19; similar to GenBank Accession Number S36895, from M. bovis	ribosomal protein L22	ribosomal protein S3 (rpS3)	ribosomal protein L16	ribosomal protein CtrL29e	ribosomal protein S17e	ribosomal protein CtrL14e	ribosomal protein CtrL24e	ribosomal protein Ctv. Se
stop	914485	915136	915467	916633	916539	917627	918304	918655	919533	919829	920157	920840	921294	921514	921758	922143	922491	923035
begin	913691	914516	915144	915629	916051	916965	917612	918323	918682	919542	919723	920184	920866	921272	921510	921778	922159	922496
ORF	ORF936	ORF937	ORF938	ORF939	ORF940	ORF941	ORF942	ORF943	ORF944	ORF945	ORF946	ORF947	ORF948	ORF949	ORF950	ORF951	ORF952	ORF953

1%	86	100	66	66	66	66	100	86	97	86	88					14	38	09
Score	487	927	605	814	740	2254	604	646	847	1040	1735					250	258 .	264
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis					Haemophilus influenzae	Haemophilus influenzae	Helicobacter pylori
a	M80325	M60652	M80325	M80325	M80325	L25077	L33834	L33834	L33834	L33834	U83198					U32717	U32716	AE000540
Homology	ribosomal protein CtrS8e	ribosomal protein L6	ribosomal protein CtrL18e	ribosomal protein CtrS5e	ribosomal protein CtrL15e	homolog	ribosomal protein S13	ribosomal protein S11	RNA polymerase alpha-subunit	RNA polymerase alpha-subunit	glyceraldehyde-3-phosphate dehydrogenase	putative	putative	putative	putative	crossover junction endodeoxyribonuclease (ruvC)	Holliday junction DNA helicase (ruvA)	nucleoside diphosphate kinase (ndk)
stop	923453	924032	924425	924937	925364	926760	927184	927604	928155	928759	930244	930656	931078	931666	931959	932579	933201	933621
begin	923160	923484	924048	924443	924933	925390	926819	927209	927577	928100	929222	930222	930608	931367	931549	932070	932602	933319
ORF	ORF954	ORF955	ORF956	ORF957	ORF958	ORF959	ORF960	ORF961	ORF962	ORF963	ORF964	ORF965	ORF966	ORF967	ORF968	ORF969	ORF970	ORF971

٩	7.2	\$	36	51	2	33	40	36	42	32	89	જ	44		į		42	1	F		
0.000	arose	186	156	1562	2002	120	668	265	1334	198	882	417	755		223	644	260	170	9/1		
Sheries		Myxococcus xantnus	Mycoplasma genitalium	Pseudomonas putida	Bacillus subtilie		Arabidopsis thaliana	Bacillus subtilis		Caenorhabditis elegans	Pseudomonas fluorescens	Deinococcus radiodurans	Oryza sativa		Bacillus aubtilie		Mus musculus	Cucumis sativns			
qī	TOESOS		U39706	X62540	D26185	AE000610	Z49227	AF008220	D12982	227079	L27278	127276	M31616		298682		M59288	D26106			
Homology	nucleogide 5'-dinhosphate		hypothetical protein (GB:U14003_297)	homologous to E.coli gidA	replicative DNA helicase	phosphatidylglycerophosphate synthase (pgsA)	adenine nucleotide translocase	putative protease	DNA polymerase	T05G5.5	The first ATG in the open reading frame was chosen as the initiation codon.	The first GTG in the open reading frame was chosen as the initiation codon.	ADPglucose pyrophosphorylase	putative	YlbH protein	putative	ferrochelatase	ferrochelatase	putative	putative	putative
stop	933785		933848	934539	939986	939098	940933	942068	944685	945287	946294	946676	948454	949277	949594	950676	951330	951643	952798	954264	955157
begin	933522		934546	936377	938081	938538	939329	941031	942082	944634	945287	946293	947105	948522	949277	949849	950680	951281	951788	953581	954426
ORF	ORF972		ORF973	ORF974	ORF975	ORF976	ORF977	ORF978	ORF979	ORF980	ORF981	ORF982	ORF983	ORF984	ORF985	ORF986	ORF987	ORF988	ORF989	ORF990	ORF991

OppB gene product  OppB gene product  dipeptide ABC transporter,  permease protein (dppC)  methylated DNA protein  Cysteine methyltransferase  putative  putative  putative  putative  putative  putative  putative  transcriptional activator of 2121  pilA  sensor protein  putative  transcriptional activator of 2121  pilA  sensor protein  putative  unknown  putative  unknown  putative  unknown  putative  unknown  putative  unknown  putative  putative  unknown  putative	ORF	pegin	stop	Homology	4			
957837         95912         OppB gene product         X56347         Bacillus subtilis         327           95929         961050         dipeptide ABC transporter, permease protein (dppC)         AE000548         Helicobacter pylori         263           961562         961063         methylated DNA protein         U67593         Methanococcus         109           961379         961867         putative         Permease protein methyltransferase         Z75208         Bacillus subtilis         775           964941         962875         sputative         Cerevisiae         775         775           964941         962896         putative         Cerevisiae         492           964941         962808         putative         Cerevisiae         492           964903         966193         unknown         Z48008         Saccharomyces         492           974866         putative         Cerevisiae         147         376           97867         97286         putative         D26185         Bacillus subtilis         147           97520         974546         97346         unknown         D26185         Bacillus subtilis         147           97650         97523         974586         putative		955754	957940	gene	X93084	Methanosarcina barkeri	Score 130	41 4
959299 961050   dipeptide AEC transporter,   AE000548   Helicobacter pylori   263		957837	959312	OppB gene nroduct	VCC242			
961562   961053   methylated DNA protein (dppC)   Methanococcus   109     961562   961053   methylated DNA protein (dppC)   Methanococcus   109     961575   961487   putative   245078   putative   2461941   965578   putative   2461941   965508   putative   2461941   965508   putative   2461941   965708   putative   2461941   965903   putative   2461941   965903   putative   2461941   965903   putative   2461941   965903   putative   2461941   966903   putative   2461941   966903   putative   2461941   putative   246194   putati		000000	020170	offe game produce	A3634 /	- 1	327	ဗ္က
961562         961053         methylated DNA protein         U67593         Methanococcus         109           961575         961084         putative         109         109         109           961979         961584         putative         27508         Paca subunit         775         175           964941         965108         putative         278008         Saccharomyces         492           1 967023         968064         putative         248008         Saccharomyces         492           1 967044         968061         putative         Cerevisiae         492           1 968003         968064         putative         Cerevisiae         492           1 968003         968064         putative         Apacococus xanthus         147           2 968003         968064         putative         Bened putative         147           3 97806         971064         sensor protein         D64126         Bacillus subtilis         147           976103         976104         mknown         D64126         Bacillus subtilis         141           97620         97621         mknown         Z49939         Saccharomyces         183           976580         97689         putat		66266	OCOTOR	dipeptide ABC transporter, permease protein (dppC)	AE000548	Helicobacter pylori	263	98 36
962575         961487         putative         Jannaschii           964914         962546         putative         775           964914         962548         putative         775           964914         962508         putative         775           964914         965061         putative         775           964914         965062         putative         78008           967023         966061         putative         78008           967024         966061         putative         78008           968003         968064         putative         78008           970685         968064         putative         78008           970886         971024         sensor protein         139904         Myxococcus xanthus         147           971806         971024         sensor protein         139904         Myxococcus xanthus         147           971086         971086         putative         153904         Myxococcus xanthus         147           971086         971086         putative         147         147           976103         976104         putative         142         142           976104         97688         putative		961562	961053	methylated DNA protein	U67593	Methanococcus	109	39
961979         Pottative         P				cysteine metnyitransierase		jannaschii		
961979         961584         putative         775           964914         962945         phenylalanyl-tRNA synthetase beta subunit         275208         Bacillus subtilis         775           967023         966193         unknown         248008         Saccharomyces         492           967024         968061         putative         cerevisiae         492           967025         968062         putative         cerevisiae         849           970685         969528         transcriptional activator of pilA         21154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         Li39904         Myxococcus xanthus         147           974546         973746         unknown         D64126         Bacillus subtilis         140           97520         974558         unknown         D26185         Bacillus subtilis         141           97650         97650         97650         Saccharomyces         183         3           97650         97650         97650         Saccharomyces         183         3           97650         97650         97650         Saccharomyces         183         3           97650         97650         97		962575	961487	putative				
964914         962545         phenylalanyl-tRNA synthetase         Z75208         Bacillus subtilis         775           964941         96508         putative         248008         Saccharomyces         492           1 967023         966193         unknown         248008         Saccharomyces         492           2 968903         968064         putative         cerevisiae         492           2 970885         969528         transcriptional activator of pila         212154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         L39904         Mycococcus xanthus         147           974546         971024         sensor protein         D64126         Bacillus subtilis         500           97523         974546         unknown         D26185         Bacillus subtilis         141           97623         974558         unknown         A24939         Saccharomyces         183           976520         976554         unknown         Z49339         Saccharomyces         183           976580         976886         976896         putative         249339         Saccharomyces         183           976886         977681         peptide release factor 2		961979	961584	putative				
964941         965708         putative         492           967023         966193         unknown         248008         Saccharomyces         492           967444         968061         putative         cerevisiae         492           968903         968064         putative         kanscriptional activator of pilA         212154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         L39904         Myxococcus xanthus         147           971806         971024         sensor protein         L39904         Myxococcus xanthus         147           974546         973746         unknown         D54126         Bacillus subtilis         500           97523         974558         unknown         D564126         Bacillus subtilis         141           976193         975207         hypothetical protein in htrA         AB000126         Escherichia coli         142           97658         97658         yr6899         putative         cerevisiae         183           97658         97658         yr6899         putative         cerevisiae         183           97658         97658         yr6896         putative         249939         cerevisiae		964914	962545	phenylalanyl-tRNA synthetase beta subunit		Bacillus subtilis	775	37
967023         966193         unknown         Z48008         Saccharomyces         492           967444         968061         putative         cerevisiae         492           968903         968064         putative         cerevisiae         849           9710685         969528         transcriptional activator of pilA         L139904         Myxococcus xanthus         147           971806         971024         sensor protein         L139904         Myxococcus xanthus         147           974546         973746         unknown         D64126         Bacillus subtilis         500           97523         974558         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrh         AB000126         Bscherichia coli         142           976588         976899         putative         245939         cerevisiae         183           976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977681         977833         petaleae factor 2         AF013188         Bacillus subtilis         187		964941	965708	putative				
967444         968061         putative         849           968903         968064         putative         370685         969528         transcriptional activator of pilA         212154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         L39904         Myxococcus xanthus         147           973053         972388         putative         D64126         Bacillus subtilis         141           975203         974558         unknown         D26185         Bacillus subtilis         142           976520         976254         unknown         Z49939         Saccharomyces         183           976588         976899         putative         Z49939         Cerevisiae         37           976686         977635         peptide release factor 2         X99401         Bacillus firmus         57           977661         977818         978433         putative         187         187	0.0	967023	966193	unknown	248008	Saccharomyces cerevisiae	492	4
968903         968064         putative         4           970685         transcriptional activator of pilA         12154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         Li39904         Myxococcus xanthus         147           973053         972388         putative         D64126         Bacillus subtilis         500           975223         974558         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrh         AB000126         Bscherichia coli         142           976520         976524         unknown         249939         Saccharomyces         183           976580         976886         976899         putative         Cerevisiae         183           976686         977635         peptide release factor 2         X99401         Bacillus firmus         534           977618         978413         putative         187         4	1	967444	968061	putative				
970685         969528         transcriptional activator of pilA         12154         Pseudomonas aeruginosa         849           971806         971024         sensor protein         Li39904         Myxococcus xanthus         147           973053         97238B         putative         D64126         Bacillus subtilis         500           975223         97455B         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrA         AB000126         Escherichia coli         142           976520         976520         976254         unknown         Z49939         Saccharomyces         183           976580         976886         976899         putative         X99401         Bacillus firmus         534           977661         977818         977813         Bacillus subtilis         187         4	7	806896	968064	putative				
971806         971024         sensor protein         L39904         Myxococcus xanthus         147           973053         97238B         putative         D64126         Bacillus subtilis         500           97523         97455B         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrA         AB000126         Escherichia coli         142           976520         976254         unknown         Z49939         Saccharomyces         183           97658B         97688         97689         putative         X99401         Bacillus subtilis         187           977618         977633         release factor 2         AF01318B         Bacillus subtilis         187	m .	970685	969528	scriptional activator			849	45
973053         971024         Sensor protein         L39904         Myxococcus xanthus         147           973053         972388         putative         D64126         Bacillus subtilis         500           97523         974558         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrA         AB000126         Escherichia coli         142           976520         976254         unknown         Z49939         Saccharomyces         183           976588         976896         Putative         X99401         Bacillus firmus         534           977618         977633         release factor 2         X99401         Bacillus subtilis         187		200160	100100			,		
974546         973746         unknown         D64126         Bacillus subtilis         500           97523         974558         unknown         D26185         Bacillus subtilis         141           976193         97520         hypothetical protein in htrh         AE000126         Bacillus subtilis         142           976520         976524         unknown         Z49939         Saccharomyces         183           976588         976896         putative         Cerevisiae         183           976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977933         release factor 2         AF013188         Bacillus subtilis         187	, ,	277060	\$70T/6	sensor protein	L39904		147	ထ
974540         Unknown         D64126         Bacillus subtilis         500           97523         974558         unknown         D26185         Bacillus subtilis         141           976193         975207         hypothetical protein in htrA         AB000126         Escherichia coli         142           976520         976524         unknown         249939         Saccharomyces         183           976588         976899         putative         cerevisiae         183           976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977643         putative         AF013188         Bacillus subtilis         187	,	973033	9/2388	putative				
976193         976193         975207         hypothetical protein in htrA dapD intergenic region         AE000126         Escherichia coli         141           976193         975207         hypothetical protein in htrA dapD intergenic region         249939         Saccharchia coli         142           976520         976254         unknown         Z49939         Saccharcmyces         183           976588         976896         putative         Cerevisiae         183           976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977918         978433         putative         187         187	,	97533	9/3/46	пикпомп	D64126		200	20
976520         976254         unknown         Z49939         Saccharomyces         183           976588         97689         putative         x99401         Bacillus firmus         534           977661         977633         release factor 2         X99401         Bacillus subtilis         187		975163	9/4558		D26185		141	44
976520         976254         unknown         Z49939         Saccharomyces         183           976588         976899         putative         cerevisiae         ,           976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977918         978433         putative         187	0	561976	107616		<b>AE</b> 000126	Escherichia coli	142	42
976588         976899         putative         X99401         Bacillus firmus         534           977661         977933         release factor 2         AF013188         Bacillus subtilis         187           977918         978433         putative         187	6	976520	976254	unknown	Z49939	Saccharomyces cerevisiae	183	39
976886         977635         peptide release factor 2         X99401         Bacillus firmus         534           977661         977933         release factor 2         AF013188         Bacillus subtilis         187           977918         978433         putative         187		976588	976899	putative				
977661         977933         release factor 2         AF013188         Bacillus subtilis         187           977918         978433         putative		976886	977635	,	X99401	- 1	534	
977918 978433 putative	2	977661	977933	release factor 2	AF013188		101	F 2
	8	977918	978433	putative			101	3

ORF	begin	stop	Homology	A	Species	Score	*I
ORF1035	998962	999225	9-kDa cysteine-rich outer membrane protein	M35148	Chlamydia trachomatis	504	100
ORF1036	999375	1001033	outer membrane protein 2	M23001	Chlamydia trachomatis	2857	100
ORF1037	1001211	1001516	15-kDa serine-rich outer membrane protein	M35148	Chlamydia trachomatis	276	\$
ORF1038	1001392	1001664	15-kDa serine-rich outer membrane protein	M35148	Chlamydia trachomatis	438	97
ORF1039	1003721	1001823	ORF of prc gene (alt.)	D00674	Escherichia coli	486	42
ORF1040	1004459	1004845	StrA	M86701	Haemophilus influenzae	454	92
ORF1041	1004990	1005382	ribosomal protein S7	211567	Chlamydia trachomatis	662	66
ORF1042	1005391	1007496	translation elongation factor EF-G (fusA)	AE000625	Helicobacter pylori	2147	62
ORF1043	1007486	1007821	ribosomal protein S10	221676	Spirulina platensis	350	89
ORF1044	1007802	1008698	NADPH-sulfite reducatase flavoprotein component	M23008	Escherichia coli	113	48
ORF1045	1009426	1009121	unknown	292774	Mycobacterium tuberculosis	102	42
ORF1046	1010534	1012054	serine hydroxymethyltransferase	238002	Bacillus subtilis	1021	55
ORF1047	1012397	1011942	putative				
ORF1048	1012042	1012635	ATP-dependent Clp protease proteolytic subunit	D90915	Synechocystis sp.	365	4
ORF1049	1012593	1012862	putative				
ORF1050	1012811	1013440	diaminopimelate epimerase (dapF)	U32759	Haemophilus influenzae	108	40
ORF1051	1013456	1014055	putative			•	
ORF1052	1013977	1014489	putative				

1%	38		42	!				33	36				43		8	51	37	35	40	35	49		36			46		
Score	263		428					164	201				218		251	603	161	439	312	354	95		75			160	_	
Species	Escherichia coli		Helicobacter pylori	4				Helicobacter pylori	լ թ	tuberculosis			Escherichia coli		Porphyra purpurea	Escherichia coli	Haemophilus influenzae	Synechocystis sp.	Bacillus subtilis	Synechocystis sp.	Chlamydia psittaci	1	Chlamydia psittaci			Chlamydia psittaci		
£	AE000459		AE000579					AE000647	295208				U18997		U38804	U18997	U32769	D90903	AB000617	D90903	U72499		U72499			U72499		
Homology	hypothetical 28.1 kD protein in udp-rfaH intergenic region	putative	conserved hypothetical	protein	putative	putative	putative	hemolysin	unknown		putative	putative	50S ribosomal subunit	protein L21	50S ribosomal protein L27	ORF_f390	GTP-binding protein (obg)	hypothetical protein	YcdI	adhesion protein	putative 98 kDa outer	membrane protein		membrane protein	putative		membrane protein	putative
stop	1014529	1015145	1015939		1017245	1017916	1018580	1019831	1020114		1021075	1022097	1023667		1023949	1024776	1025045	1024967	1025839	1026546	1027929		1030508		1032086	1033456		1035910
begin	1015224	1016002	1017120		1017766	1018911	1019191	1020199	1021007		1021569	1022411	1023344		1023701	1023976	1024704	1025881	1026546	1027379	1030604		1033252		1031733	1037037		1035674
ORF	ORF1053	ORF1054	ORF1055		ORF1056	ORF1057	ORF1058	ORF1059	ORF1060		ORF1061	ORF1062	ORF1063		ORF1064	ORF1065	ORF1066	ORF1067	ORF1068	ORF1069	ORF1070		ORF1071		ORF1072	ORF1073		ORF1074

Species Score T&	
B	
Homology	putative
gcop	1036501
begin	1036175
ORF	ORF1075

200	negin	втор	Homology	ΙD	Species	Score	%I
ORF1076	68 (com)	1036967	putative				
ORF1077	16591	16989	GutQ/KpsF Family Sugar-P Isomerase	AE001313	Chlamydia trachomatis	658	97
ORF1078	31779	31408	putative				
ORF1079	56502	56834	hypothetical protein	AE001309	Chlamydia trachomatis	284	95
ORF1080	26686	56913	hypothetical protein	AE001309	Chlamydia trachomatis	303	94
ORF1081	64748	65074	hypothetical protein (possible 357R?)	AE001309	Chlamydia trachomatis	501	100
ORF1082	73482	73195	Predicted OMP [leader (19) peptide]	AE001308	Chlamydia trachomatis	476	100
ORF1083	78482	78736	putative				
ORF1084	79803	79411	hypothetical protein	AE001307	Chlamydia trachomatis	583	86
ORF1085	82333	81959	Lon ATP-dependent protease	AE001307	Chlamydia trachomatis	607	66
ORF1086	87313	86999	hypothetical protein	AE001307	Chlamydia trachomatis	534	100
ORF1087	109929	109456	hypothetical protein	AE001305	Chlamydia trachomatis	529	98
ORF1088	111599	111351	putative				
ORF1089	112069	111734	putative				
ORF1090	112666	112911	hypothetical protein	AE001305	Chlamydia trachomatis	395	94
ORF1091	114017	113715	putative				
ORF1092	120757	120464	putative				
ORF1093	125133	125522	predied ferredoxin	AE001303	Chlamydia trachomatis	631	97
ORF1094	131888	131604	putative			•	T
ORF1095	144164	144427	putative				
ORF1096	150698	150369	putative				

ORF	begin	stop	Homology	CI CI	Sperion	Santa	į
ORF1097	164385	163948	NADH (Ubiquinone) Dehydrogenase	AE001300	Chlamydia trachomatis	755	100
ORF1098	165690	166115	hypothetical protein	AE001300	Chlamydia trachomatis	724	66
ORF1099	168742	168425	hypothetical protein	AE001300	Chlamydia trachomatis	356	96
ORF1100	170509	170793	hypothetical protein	AE001300	Chlamydia trachomatis	489	100
ORF1101	177145	177474	AcCoA Carboxylase/Transferase Alpha	AE001299	Chlamydia trachomatis	518	66
ORF1102	188295	188023	! 1	AE001298	Chlamydia trachomatis	451	100
ORF1103	188791	188330	hypothetical protein	AE001298	Chlamydia trachomatis	733	97
ORF1104	190629	190336	putative				
ORF1105	197313	197083	putative				
ORF1106	210914	211384	putative				
ORF1107	235160	234852	Glutamate Aminomutase	AE001295	Chlamydia trachomatis	507	26
ORF1108	237227	236913	putative				
ORF1109	249733	249446	Oligopeptide Permease	AE001293	Chlamydia trachomatis	512	100
ORF1110	253493	253158	hypothetical protein	AE001293	Chlamydia trachomatis	318	63
ORF1111	253701	254789	hypothetical protein	AE001293	Chlamydia trachomatis	1860	99
ORF1112	271633	271932	hypothetical protein	AE001291	Chlamydia trachomatis	512	100
ORF1113	275666	276070	Disulfide bond Oxidoreductase	AE001291	Chlamydia trachomatis	700	66
ORF1114	277931	278218	putative				

Score I%	<del>-</del>	433 95		422 95			338 99											
Sc	-			-		+												
Species	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis		Chlamydia trachomatis		Chlamydia trachomatis			1 1 1						Chlamydia trachomatis	Chlamydia trachomatis
a	AE001290 (	AE001289 (		AE001287 C		AE001287 C		AE001287 C										
Homology	hypothetical protein	Phospholipase D Endonuclease Superfamily	putative	hypothetical protein	putative	hypothetical protein	ı	nypoumetical procein	procein protein	protein protein se	rotein rotein	rotein rotein Superfamily sptide}	rotein rotein Superfamily eptide}	rotein  Superfamily  sptide}	cal protein  idase  cal protein  ase D Superfamily  33) peptide}  cal protein  cal protein	ical protein ical protein ical protein pase D Superfamily (33) peptide} ical protein r Secretion Protein	cal protein idase  cal protein ase D Superfamily 33) peptide}  cal protein  cal protein  cal protein  sal protein  Secretion Protein	cal protein idase  cal protein ase D Superfamily 33) peptide}  cal protein cal protein  Secretion Protein al protein al protein
stop	282481	293489	303469	308965	312536	312602	312772	 : : : :	320598	320598	320598 340503 353324	320598 340503 353324 373699	320598 340503 353324 373699	320598 340503 353324 373699 377756				
Degin	282741	293178	303155	309297	312219	312853	313167		320224	320224	320224 340249 352839	320224 340249 352839 373475	320224 340249 352839 373475	320224 340249 352839 373475 377316	320224 340249 352839 373475 377316 379268	320224 340249 352839 373475 377316 379268 395098	320224 340249 352839 373475 377316 379268 395098 401594	320224 340249 352839 373475 377316 379268 395098 401594 411425
ORF	ORF1115	ORF1116	ORF1117	ORF1118	ORF1119	ORF1120	ORF1121	-	ORF1122	ORF1122 ORF1123	ORF1122 ORF1123 ORF1124	ORF1122 ORF1123 ORF1124	ORF1122 ORF1123 ORF1124 ORF1125	ORF1122 ORF1124 ORF1125 ORF1126 ORF1127	ORF1122 ORF1124 ORF1125 ORF1126 ORF1127	ORF1122  ORF1124  ORF1125  ORF1126  ORF1127  ORF1129	ORF1122 ORF1124 ORF1125 ORF1126 ORF1127 ORF1129 ORF1129	ORF1122 ORF1124 ORF1125 ORF1126 ORF1127 ORF1129 ORF1130

ORF	pegin	stop	Homology	A	Species	Score	18
ORF1133	422889	423212	Glycogen Hydrolase (debranching)	AE001278	Chlamydia trachomatis	206	100
ORF1134	427842	428183	hypothetical protein	AE001278	Chlamydia trachomatis	610	100
ORF1135	428732	429451	hypothetical protein	AE001278	Chlamydia trachomatis	1010	98
ORF1136	442557	442799	hypothetical protein	AE001277	Chlamydia trachomatis	649	94
ORF1137	443628	444041	L31 Ribosomal Protein	AE001277	Chlamydia trachomatis	538	96
ORF1138	443678	443166	putative				T
ORF1139	445901	446155	putative				
ORF1140	467981	468262	putative				
ORF1141	471869	472108	Putative Outer Membrane Protein I	AE001361	Chlamydia trachomatis	370	100
ORF1142	488032	488337	Membrane Thiol Protease	AE001360	Chlamydia trachomatis	483	96
ORF1143	497179	497694	Low Calcium Response Protein H	AE001359	Chlamydia trachomatis	864	95
ORF1144	500474	500202	putative				
ORF1145	508968	509561	ABC transporter permease	AE001358	Chlamydia trachomatis	964	100
ORF1146	510845	511264	hypothetical protein	AE001358	Chlamydia trachomatis	360	89
ORF1147	526525	526848	hypothetical protein	AE001356	Chlamydia trachomatis	242	81
ORF1148	531318	531863	hypothetical protein	AE001356	Chlamydia trachomatis	127	100
ORF1149	556826	557224	hypothetical protein	AE001354	Chlamydia trachomatis	683	66
ORF1150	564971	564537	hypothetical protein	AE001353	Chlamydia trachomatis	534	100

%i	53	100	99	97	100	98		66					66			86				98		96
Score	220	925	441	273	442	1176		434					324			864				606	-	661
Species	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis					Chlamydia trachomatis			Chlamydia trachomatis				Chlamydia trachomatis		Chlamydia trachomatis
ET.	AE001353	AE001353	AE001353	AE001352	AE001351	AE001350		AE001349					AE001345			AB001343				AE001336		AE001335
Homology	Glycerol-3-P Acyltransferase	Insulinase family/Protease III	hypothetical protein	General Stress Protein	hypothetical protein	hypothetical protein	putative	hydrolase/phosphatase homolog	putative	putative	putative	putative	Phenolhydrolase/NADH ubiquinone oxidoreductase	putative	putative	hypothetical protein	putative	putative	putative	(Pseudouridine Synthase)	putative	hypothetical protein
stop	567232	570890	571332	575801	590650	598593	606626	607786	610391	633353	637482	649924	652562	655325	660810	677057	679253	732210	742383	758782	760521	770894
ujbeq	566963	570351	571072	576025	590363	597868	688909	608031	610110	632703	637213	650517	652317	654753	661118	677596	679528	732536	742069	759318	760282	771313
ORF	ORF1151	ORF1152	ORF1153	ORF1154	ORF1155	ORF1156	ORF1157	ORF1158	ORF1159	ORF1160	ORF1161	ORF1162	ORF1163	ORF1164	ORF1165	ORF1166	ORF1167	ORF1168	ORF1169	ORF1170	ORF1171	ORF1172

18	66					98	66		95				97		66	100	98		97		98	
Score	520					268	747		551				195		410	593	542		467		647	
Species	Chlamydia trachomatis					Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis				Chlamydia trachomatis		Chlamydia trachomatis	Chlamydia trachomatis	Chlamydia trachomatis		Chlamydia trachomatis		Chlamydia trachomatis	
A	AE001335					AE001327	AE001327		AE001326				AE001324		AE001322	AE001322	AE001321		AE001320		AE001320	
Homology	hypothetical protein	putative	putative	putative	putative	hypothetical protein	hypothetical protein	putative	hypothetical protein	putative	putative	putative	Myristoyl GlcNac Deacetylase	putative	hypothetical protein	hypothetical protein	hypothetical protein	putative	2-Component Sensor	putative	Phosphoglycolate Phosphatase	putative
stop	772408	788457	815967	846914	868054	875658	876915	884312	891467	900417	902269	907783	912567	935741	946692	952783	965873	968765	970731	972404	973508	998404
pegin	772115	788137	816302	846606	867803	875386	876445	884548	891859	900770	902553	908046	912313	935451	946961	953193	966199	969298	971009	972162	973119	998649
ORF	ORF1173	ORF1174	ORF1175	ORF1176	ORF1177	ORF1178	ORF1179	ORF1180	ORF1181	ORF1182	ORF1183	ORF1184	ORF1185	ORF1186	ORF1187	ORF1188	ORF1189	ORF1190	ORF1191	ORF1192	ORF1193	ORF1194

t t							
OKE	Degin	Stop	Homology	#	Species	04000	40
						0 100	e T
ORF1195	1004280	1003882	hypothetical protein	AE001317	Chlamydia trachomatis	571	99
			hamon hand and				
ORF1196	1010200	1009532	Mypochecical procein	AE001317	Chlamydia trachomatis	1132	66
					-		
201120	100001	20000					
ORFILSI	10231/4	1029482	putative				

TABLE 2

SEQ ID NO	begin	stop	preferred star
2	501	208	501
3	3276	505	3153
4	5068	3242	5062
5	6400	5126	6400
6	7977	6619	7977
7	8582	8082	8582
8	8995	8591	8995
9	9440	8979	9440
10	9828	10430	9828
11	10367	11254	10430
12	11245	11916	11245
13	12068	13324	12068
14	13532	14413	13538
15	14807	15019	14807
16	14932	15969	14977
17	15995	16501	16004
18	16467	16138	16377
19	18190	17417	18178
20	20521	18437	20518
21	22202	20814	22166
22	22602	22153	22509
23	22804	22478	22795
24	23183	22824	23180
25	23394	23110	23394
26	24569	23394	24569
27	26383	24641	26383
28	26640	27710	26640
29	28780	27725	28729
30	29957	28740	29957
31	30721	30032	30628
32	31281	30520	31254
33	31436	31780	31436

SEQ ID NO	begin	stop	preferred start
34	33356	31800	33344
35	33901	33314	33874
36	34116	35027	34116
37	34988	35359	35027
38	35167	35919	35377
39	35923	36996	36031
40	37810	37013	37765
41	38207	39085	38252
42	39151	39927	39157
43	39923	40756	39959
44	40760	42007	40772
45	42175	43116	42229
46	42999	43802	43128
47	44211	45227	44217
48	46072	45275	46066
49	46340	45975	46331
50	46895	46506	46865
51	47955	46882	47955
52	48585	48178	48558
53	50072	48630	50012
54	50710	50099	50692
55	52439	50925	52430
56	53484	52348	53478
57	54536	53466	54536
58	55086	54595	55104
59	56350	55031	56350
60	55659	56084	55722
61	56847	58235	56931
62	58423	59181	58423
63	59185	60195	59194
64	60188	61483	60191
65	61496	62353	61496
66	62500	63141	62518
67	63396	63983	63396

SEQ ID NO	begin	stop	preferred star
68	64628	64071	64580
69	64285	64656	64285
70	64944	64609	64938
71	65347	67269	65347
72	67656	68873	67815
73	68877	69233	68892
74	69212	69721	69323
75	69958	70455	69970
76	70701	71006	70725
77	73191	71086	73185
78	74900	73497	74891
79	75463	74876	75463
80	77124	75502	77124
81	77000	77299	77012
82	78095	77145	78095
83	79065	78154	79065
84	81971	79878	81965
85	82639	83271	82642
86	83792	84850	83921
87	84876	86921	84888
88	88650	87313	88383
89	87440	87805	87458
90	88400	88747	88409
91	88717	89265	88729
92	89355	89732	89355
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96	93138	92344	92874
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107	105259	105894	105271
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112	111330	112520	111330
113	112915	113463	112918
114	113566	113994	113566
115	114020	114604	114020
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119	119823	124010	119823
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122	126261	125536	126243
123	126328	126930	126331
124	127138	127785	127147
125	127924	129714	127942
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128	131834	133156	131852
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130	133625	133999	133628
131	133861	134508	133948
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142	148115	148549	148115
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144	149000	149305	149033
145	149187	149708	149187
146	149712	150911	149769
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162	166686	166093	166686
163	168171	166729	168171
164	169249	168848	169189
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175	180704	180048	180635
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181	186848	186000	186791
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183	187426	187809	187429
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185	189693	190352	189693
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187	190785	191786	190824
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197	200327	199941	200306
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213	215143	214754	215128
214	216705	215236	216705
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228	225334	225549	225328
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230	227299	226769	227170
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232	228457	227750	228439
233	230001	228607	229854
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243	242022	240313	242022
244	242846	241941	242846
245	244480	242798	244456
246	245897	244479	245891
247	246877	245924	246829
248	247731	246985	247725
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251	250383	249766	250383
252	251186	250545	251174
253	252111	251095	252099
254	253088	252066	253088
255	255153	256718	255153
256	256762	257844	256774
257	257911	258690	257962
258	258780	259187	258840
259	259193	261604	259193
260	261622	264129	261622
261	264125	264742	264134
262	264741	265628	264759
263	266416	265631	266416
264	266938	266426	267946
265	267961	266942	267946
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267	268510	268205	268510
268	270116	268500	270116
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277	279771	279013	279870
278	280777	279767	280762
279	281603	281295	281576
280	282104	281787	282086
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283	284817	285674	284844
284	285637	286137	285670
285	286357	286677	286399
286	286681	287898	286852
287	288127	289227	288358
288	289744	290679	289744
289	290828	291535	291206
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291	292326	293048	292350
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293	295684	295010	295684
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297	298905	300458	298920
298	302152	300527	302131
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300	306157	304973	306142
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311	316507	315665	316507
312	317284	316529	317284
313	317592	317338	317592
314	318470	317499	318416
315	317599	317874	317599
316	318947	318477	318887
317	319342	320142	319342
318	320544	321497	320682
319	321485	321937	321497
320	321901	322362	321943
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322	323144	324913	323177
323	325621	324977	325621
324	326268	325621	326262
325	326469	327203	326469
326	327281	328150	327302
327	328605	328204	328602
328	329066	328734	329066
329	329663	329292	329648
330	330666	329608	330663
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333	332404	331721	332404
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338	335142	334729	335124
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345	341783	342022	341786
346	342249	342470	342249
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348	343361	344032	343379
349	343956	344225	343962
350	344357	345142	344357
351	345934	345161	345934
352	347102	346080	347102
353	347113	347940	347119
354	350164	348146	350113
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356	352207	351314	352207
357	352727	352245	352703
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360	354721	354140	354721
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362	356700	357377	356700
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365	360753	361121	360753
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368	363859	362816	363859
369	364116	365195	364116
370	365198	365587	365198
371	365479	367320	365614
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374	369088	370251	369088
375	370769	371086	370769
376	371203	372816	371209
377	373119	373529	373152
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381	377062	376748	377038
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393	386127	386627	386232
394	387372	386872	387351
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406	398753	397401	398747
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414	405165	405920	405165
415	407049	405955	407049
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418	411707	413410	411722
419	413433	412606	413334
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421	413841	415112	413991
422	414379	413978	414220
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425	418053	417721	418044
426	418603	418031	418582
427	419531	418647	419531
428	420190	419672	420190
429	421171	420245	421171
430	421988	421518	421988
431	422486	423043	422492
432	423226	425079	423295
433	426054	425146	426021
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435	427248	427817	427248
436	429560	429886	429623
437	430360	429857	430360
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439	430933	431787	430966
440	431658	431987	431688
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445	438262	437894	438256
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447	439339	438986	439339
448	439705	439358	439705
449	441042	439699	441042
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451	442593	441898	442584
452	444505	446388	444505
453	448068	446452	448029
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455	450546	451076	450576
456	451623	451144	451401
457	452593	451517	452575
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459	453567	454868	453567
460	455430	454972	455403
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462	457384	456047	457384
463	457659	458450	457659
464	458508	459632	458511
465	459839	461203	459839
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467	461887	462621	462151
468	463758	462895	463749
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471	467420	466113	467414
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480	475198	474602	475195
481	476527	475613	476509
482	478640	476517	478640
483	479084	478665	479078
484	479723	479088	479720
485	480012	479668	479898
486	481466	479895	481412
487	481732	481496	481732
488	481864	483429	481870
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490	484898	487864	484970
491	485725	485222	485593
492	488204	489247	488321
493	488571	488233	488562
494	489440	490456	489473
495	492765	490507	492690
496	492357	492893	492654
497	493744	492737	493723
498	493875	494675	493875
499	494573	494869	494609
500	494835	495365	494955
501	495174	494872	495174
502	495687	496634	495732
503	496295	497176	496523
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505	498280	499239	498301
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516	509600	510691	509600
517	511039	511527	511147
518	511547	512185	511547
519	512382	513092	512385
520	514287	513055	514269
521	514789	515244	514792
522	514994	515269	515027
523	515553	515804	515553
524	515808	516422	515820
525	516476	517171	516605
526	517927	517400	517927
527	518096	518380	518114
528	518403	518822	518412
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530	519577	520497	519730
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534	524591	523623	524510
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536	525731	526078	525752
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538	526301	526735	526361
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552	540906	541805	541002
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556	544762	544487	544747
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562	548692	549459	548704
563	550385	549663	550376
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565	553041	551797	553041
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567	556300	554927	556300
568	556524	556904	556524
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570	557810	558235	557810
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573	562931	561150	562898
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575	563593	563943	563644
576	565379	566953	565379
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582	572879	573427	573077
583	574160	573660	574160
584	574426	574184	574426
585	574781	574446	574781
586	575243	574923	575156
587	575458	575057	575458
588	575849	575469	575735
589	576545	578023	576545
590	578673	578017	578673
591	579012	582104	579012
592	582697	582206	582682
593	583122	582811	583095
594	583514	583182	583484
595	583869	583438	583803
596	584435	583827	584399
597	584967	584299	584967
598	585297	585016	585285
599	585240	586610	585300
600	586484	587758	586505
601	587786	589408	587786
602	589198	589578	589258
603	590061	589630	589971
604	590739	591272	590775
605	592406	592765	592406
606	593145	592849	593127
607	593900	593121	593894
608	594138	595637	594138
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610	596864	596154	596828
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SEQ ID NO	begin	stop	preferred start
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613	601876	600734	601864
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617	604549	605610	604549
618	606619	605582	606619
619	606843	607493	606867
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621	609652	609296	609652
622	611860	610109	611830
623	611812	612927	611815
624	613597	612938	613444
625	613952	613692	613952
626	614315	615244	614441
627	615396	615683	615396
628	617711	615864	617624
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631	620416	619247	620401
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633	621184	620420	621154
634	621690	621154	621678
635	622399	621674	622399
636	623466	622414	623421
637	624178	623570	624106
638	624918	624073	624918
639	625346	626665	625367
640	626514	626900	626652
641	626954	627853	626984
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643	628715	628146	628715
644	628932	629801	628935
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648	637488	638084	637488
649	638036	640207	638111
650	640221	643472	640236
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655	645840	647111	645840
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661	652799	653593	652892
662	659884	661851	660136
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671	668121	668375	668139
672	668470	668174	668404
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674	669892	669485	669892
675	670780	669998	670765
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685	679342	680502	679342
686	680579	681280	680654
687	681539	682558	681557
688	682554	683087	682578
689	683164	684465	683164
690	684774	684418	684639
691	684839	686203	684839
692	686197	687204	686203
693	687341	688360	687341
694	688432	688193	688426
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696	689960	689631	689939
697	690487	689846	690445
698	690717	690463	690717
699	691871	690672	691856
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703	698084	697170	697958
704	698392	697979	698380
705	698792	700117	698792
706	700269	700895	700269
707	700912	702165	700990
708	702183	703412	702183
709	703522	705000	703531
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720	714232	713651	714217
721	714632	714120	714617
722	715592	714834	715739
723	715854	715558	715854
724	716937	715921	716886
725	718357	717149	718357
726	718500	718862	718590
727	719797	718499	719776
728	720273	719782	720147
729	720452	720144	720452
730	720613	721575	720613
731	721559	722356	721571
732	723248	722397	723239
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734	725763	724576	725760
735	726519	725767	726519
736	726819	726538	726801
737	727493	726753	727466
738	727984	727469	727984
739	728778	728329	728718
740	729346	728759	729334
741	732639	729442	732639
742	733246	734427	733246
743	734814	735659	734814
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752	744298	743306	744292
753	744714	744430	744660
754	744985	744611	744931
755	745557	744958	745548
756	746412	745561	746409
757	746772	746416	746697
758	748269	746944	748269
759	748966	748274	748954
760	749426	748965	749411
761	749702	749433	749681
762	750029	749721	750020
763	752307	750007	752307
764	752913	752503	752901
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766	755000	756814	755000
767	756796	758301	756832
768	758691	758446	758688
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770	760242	759871	760188
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773	761759	762142	761759
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776	764857	764438	764821
777	766068	764821	765972
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786	772390	773391	772390
787	774221	773427	774215
788	776035	774191	776035
789	776663	777706	776894
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791	779222	777732	779180
792	779321	781552	779360
793	781297	782442	781351
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796	786580	785546	786580
797	787741	786611	787729
798	787620	788021	787782
799	790124	787920	790064
800	790160	790609	790178
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807	796278	797015	796311
808	796979	797365	796979
809	797260	797856	797395
810	797772	798086	797805
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812	798925	798416	798916
813	799301	799927	799301
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825	811725	812342	811824
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827	813455	813772	813455
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829	815213	814314	815207
830	814878	814396	814975
831	815733	815428	815733
832	816116	817456	816170
833	817608	819320	817608
834	819324	819713	819342
835	819704	820402	819713
836	820375	821061	820453
837	821043	821537	821043
838	821646	822239	821667
839	822182	822931	822221
840	824355	823045	824352
841	825894	824359	825891
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843	826340	827026	826340
844	827014	827250	827014
845	827856	827230	827856
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855	835778	835371	835775
856	836482	835775	836470
857	836602	837264	836617
858	837209	838699	837209
859	838760	839575	838760
860	839942	840583	839951
861	840445	841713	840451
862	841659	842459	841686
863	842523	843068	842541
864	843495	843031	843447
865	843239	846196	843335
866	844137	843802	844077
867	848043	846217	848022
868	850123	848150	850099
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872	855525	854920	855468
873	856240	855437	856240
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875	859439	857451	859430
876	859946	859587	859916
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888	869270	871372	869336
889	871299	872582	871359
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891	872773	873915	872773
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893	874028	874438	874067
894	874778	875386	874796
895	875774	876382	875843
896	877872	877000	877866
897	878172	877876	878157
898	879098	878172	879098
899	878883	879161	878886
900	879842	879105	879809
901	880885	880052	880885
902	881863	880889	881863
903	882904	881948	882901
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906	884996	884508	884984
907	888777	885166	888771
908	890172	888940	890172
909	891164	890325	891146
910	891463	891116	891427
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923	905725	906474	905725
924	906493	906945	906493
925	907306	907001	907306
926	908101	908742	908131
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930	910081	910569	910090
931	910615	910944	910636
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941	916965	917627	916965
942	917612	918304	917612
943	918323	918655	918323
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956	924048	924425	924057
957	924443	924937	924443
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962	927577	928155	927577
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973	934546	933848	934546
974	936377	934539	936377
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976	938538	939098	938595
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978	941031	942068	941076
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980	944634	945287	944673
981	945287	946294	945287
982	946293	946676	946368
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989	951788	952798	951803
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991	954426	955157	954429
992	955754	957940	955766
993	957837	959312	957867
994	959299	961050	959317
995	961562	961053	961562
996	962575	961487	962545
997	961979	961584	961979
998	964914	962545	964914
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1001	967444	968061	967459
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1004	971806	971024	971785
1005	973053	972388	973026
1006	974546	973746	974546
1007	975223	974558	975214
1008	976193	975207	976193
1009	976520	976254	976511
1010	976588	976899	976588
1011	976886	977635	976934
1012	977661	977933	977682
1013	977918	978433	977933
1014	978619	978984	978619
1015	978933	979331	978987
1016	981197	979389	981197
1017	979711	980112	979753
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1023	986693	986046	986693
1024	987607	986693	987607
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1028	989693	993442	989693
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1077	16591	16989	16597
1078	31779	31408	31764
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1084	79803	79411	79773
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1120 312853 312602 312844		309297	308965	309297
		312219	312536	312246
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1135         428732         429451         428732           1136         442557         442799         442524           1137         443628         444041         443628           1138         443678         443166         443678           1139         445901         446155         445901           1140         467981         468262         468023           1141         471869         472108         471869           1142         488032         488337         488044           1143         497179         497694         497101           1144         500474         500202         500471           1145         508968         509561         508968           1146         510845         511264         510845           1147         526525         526848         526525           1148         531318         531863         531444           1149         556826         557224         556826           1150         564971         564537         564971           1151         566963         570351         570890         570351           1153         571072         571332         571072 </td <td>1133</td> <td>422889</td> <td>423212</td> <td>422964</td>	1133	422889	423212	422964
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1139       445901       446155       445901         1140       467981       468262       468023         1141       471869       472108       471869         1142       488032       488337       488044         1143       497179       497694       497101         1144       500474       500202       500471         1145       508968       509561       508968         1146       510845       511264       510845         1147       526525       526848       526525         1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1137	443628	444041	443628
1140       467981       468262       468023         1141       471869       472108       471869         1142       488032       488337       488044         1143       497179       497694       497101         1144       500474       500202       500471         1145       508968       509561       508968         1146       510845       511264       510845         1147       526525       526848       526525         1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1138	443678	443166	443678
1141       471869       472108       471869         1142       488032       488337       488044         1143       497179       497694       497101         1144       500474       500202       500471         1145       508968       509561       508968         1146       510845       511264       510845         1147       526525       526848       526525         1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1139	445901	446155	445901
1142       488032       488337       488044         1143       497179       497694       497101         1144       500474       500202       500471         1145       508968       509561       508968         1146       510845       511264       510845         1147       526525       526848       526525         1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1140	467981	468262	468023
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1146         510845         511264         510845           1147         526525         526848         526525           1148         531318         531863         531444           1149         556826         557224         556826           1150         564971         564537         564971           1151         566963         567232         566963           1152         570351         570890         570351           1153         571072         571332         571072           1154         576025         575801         576025	1144	500474	500202	500471
1147       526525       526848       526525         1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1145	508968	509561	508968
1148       531318       531863       531444         1149       556826       557224       556826         1150       564971       564537       564971         1151       566963       567232       566963         1152       570351       570890       570351         1153       571072       571332       571072         1154       576025       575801       576025	1146	510845	511264	510845
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1151     566963     567232     566963       1152     570351     570890     570351       1153     571072     571332     571072       1154     576025     575801     576025	1149	556826	557224	556826
1152     570351     570890     570351       1153     571072     571332     571072       1154     576025     575801     576025	1150	564971	564537	564971
1153         571072         571332         571072           1154         576025         575801         576025	1151	566963	567232	566963
1154 576025 575801 576025	1152	570351	570890	570351
	1153	571072	571332	571072
1155 590363 590650 590363	1154	576025	575801	576025
	1155	590363	590650	590363

SEQ ID NO	begin	Stop	preferred start
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1157	606889	606626	606889
1158	608031	607786	608031
1159	610110	610391	610143
1160	632703	633353	632703
1161	637213	637482	637255
1162	650517	649924	650517
1163	652317	652562	652317
1164	654753	655325	654753
1165	661118	660810	661118
1166	677596	677057	677578
1167	679528	679253	679477
1168	732536	732210	732536
1169	742069	742383	742069
1170	759318	758782	759318
1171	760282	760521	760282
1172	771313	770894	771391
1173	772115	772408	772115
1174	788137	788457	788137
1175	816302	815967	816302
1176	846606	846914	846612
1177	867803	868054	867806
1178	875386	875658	875395
1179	876445	876915	876445
1180	884548	884312	884548
1181	891859	891467	891859
1182	900770	900417	900728
1183	902553	902269	902529
1184	908046	907783	908007
1185	912313	912567	912313
1186	935451	935741	935451
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1189	966199	965873	966184

SEQ ID NO	begin	stop	preferred start
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1192	972162	972404	972165
1193	973119	973508	973119
1194	998649	998404	998625
1195	1004280	1003882	1004280
1196	1010200	1009532	1010200
1197	1029174	1029482	1029180

**TABLE 4** 

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
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5	1205	1204	3597	3596
6	1207	1206	3599	3598
7	1209	1208	3601	3600
8	1211	1210	3603	3602
9	1213	1212	3605	3604
10	1215	1214	3607	3606
11	1217	1216	3609	3608
12	1219	1218	3611	3610
13	1221	1220	3613	3612
14	1223	1222	3615	3614
15	1225	1224	3617	3616
16	1227	1226	3619	3618
17	1229	1228	3621	3620
18	1231	1230	3623	3622
19	1233	1232	3625	3624
20	1235	1234	3627	3626
21	1237	1236	3629	3628
22	1239	1238	3631	3630
23	1241	1240	3633	3632
24	1243	1242	3635	3634
25	1245	1244	3637	3636
26	1247	1246	3639	3638
27	1249	1248	3641	3640
28	1251	1250	3643	3642
29	1253	1252	3645	3644
30	1255	1254	3647	3646
31	1257	1256	3649	3648
32	1259	1258	3651	3650
33	1261	1260	3653	3652

ORFGenset	ORFoligosFd	ORFoligosFp	ORFöligosBd	ORFoligosBp
34	1263	1262	3655	3654
35	1265	1264	3657	3656
36	1267	1266	3659	3658
37	1269	1268	3661	3660
38	1271	1270	3663	3662
39	1273	1272	3665	3664
40	1275	1274	3667	3666
41	1277	1276	3669	3668
42	1279	1278	3671	3670
43	1281	1280	3673	3672
44	1283	1282	3675	3674
45	1285	1284	3677	3676
46	1287	1286	3679	3678
47	1289	1288	3681	3680
48	1291	1290	3683	3682
49	1293	1292	3685	3684
50	1295	1294	3687	3686
51	1297	1296	3689	3688
52	1299	1298	3691	3690
53	1301	1300	3693	3692
54	1303	1302	3695	3694
55	1305	1304	3697	3696
56	1307	1306	3699	3698
57	1309	1308	3701	3700
58	1311	1310	3703	3702
59	1313	1312	3705	3704
60	1315	1314	3707	3706
61	1317	1316	3709	3708
62	1319	1318	3711	3710
63	1321	1320	3713	3712
64	1323	1322	3715	3714
65	1325	1324	3717	3716
66	1327	1326	3719	3718
67	1329	1328	3721	3720

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
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69	1333	1332	3725	3724
70	1335	1334	3727	3726
71	1337	1336	3729	3728
72	1339	1338	3731	3730
73	1341	1340	3733	3732
74	1343	1342	3735	3734
75	1345	1344	3737	3736
76	1347	1346	3739	3738
77	1349	1348	3741	3740
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84	1363	1362	3755	3754
85	1365	1364	3757	3756
86	1367	1366	3759	3758
87	1369	1368	3761	3760
88	1371	1370	3763	3762
89	1373	1372	3765	3764
90	1375	1374	3767	3766
91	1377	1376	3769	3768
92	1379	1378	3771	3770
93	1381	1380	3773	3772
94	1383	1382	3775	3774
95	1385	1384	3777	3776
96	1387	1386	3779	3778
97	1389	1388	3781	3780
98	1391	1390	3783	3782
99	1393	1392	3785	3784
100	1395	1394	3787	3786
101	1397	1396	3789	3788

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
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103	1401	1400	3793	3792
104	1403	1402	3795	3794
105	1405	1404	3797	3796
106	1407	1406	3799	3798
107	1409	1408	3801	3800
108	1411	1410	3803	3802
109	1413	1412	3805	3804
110	1415	1414	3807	3806
111	1417	1416	3809	3808
112	1419	1418	3811	3810
113	1421	1420	3813	3812
114	1423	1422	3815	3814
115	1425	1424	3817	3816
116	1427	1426	3819	3818
117	1429	1428	3821	3820
118	1431	1430	3823	3822
119	1433	1432	3825	3824
120	1435	1434	3827	3826
121	1437	1436	3829	3828
122	1439	1438	3831	3830
123	1441	1440	3833	3832
124	1443	1442	3835	3834
125	1445	1444	3837	3836
126	1447	1446	3839	3838
127	1449	1448	3841	3840
128	1451	1450	3843	3842
129	1453	1452	3845	3844
130	1455	1454	3847	3846
131	1457	1456	3849	3848
132	1459	1458	3851	3850
133	1461	1460	3853	3852
134	1463	1462	3855	3854
135	1465	1464	3857	3856

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
136	1467	1466	3859	3858
137	1469	1468	3861	3860
138	1471	1470	3863	3862
139	1473	1472	3865	3864
140	1475	1474	3867	3866
141	1477	1476	3869	3868
142	1479	1478	3871	3870
143	1481	1480	3873	3872
144	1483	1482	3875	3874
145	1485	1484	3877	3876
146	1487	1486	3879	3878
147	1489	1488	3881	3880
148	1491	1490	3883	3882
149	1493	1492	3885	3884
150	1495	1494	3887	3886
151	1497	1496	3889	3888
152	1499	1498	3891	3890
153	1501	1500	3893	3892
154	1503	1502	3895	3894
155	1505	1504	3897	3896
156	1507	1506	3899	3898
157	1509	1508	3901	3900
158	1511	1510	3903	3902
159	1513	1512	3905	3904
160	1515	1514	3907	3906
161	1517	1516	3909	3908
162	1519	1518	3911	3910
163	1521	1520	3913	3912
164	1523	1522	3915	3914
165	1525	1524	3917	3916
166	1527	1526	3919	3918
167	1529	1528	3921	3920
168	1531	1530	3923	3922
169	1533	1532	3925	3924

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
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171	1537	1536	3929	3928
172	1539	1538	3931	3930
173	1541	1540	3933	3932
174	1543	1542	3935	3934
175	1545	1544	3937	3936
176	1547	1546	3939	3938
177	1549	1548	3941	3940
178	1551	1550	3943	3942
179	1553	1552	3945	3944
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181	1557	1556	3949	3948
182	1559	1558	3951	3950
183	1561	1560	3953	3952
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185	1565	1564	3957	3956
186	1567	1566	3959	3958
187	1569	1568	3961	3960
188	1571	1570	3963	3962
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190	1575	1574	3967	3966
191	1577	1576	3969	3968
192	1579	1578	3971	3970
193	1581	1580	3973	3972
194	1583	1582	3975	3974
195	1585	1584	3977	3976
196	1587	1586	3979	3978
197	1589	1588	3981	3980
198	1591	1590	3983	3982
199	1593	1592	3985	3984
200	1595	1594	3987	3986
201	1597	1596	3989	. 3988
202	1599	1598	3991	3990
203	1601	1600	3993	3992

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
204	1603	1602	3995	3994
205	1605	1604	3997	3996
206	1607	1606	3999	3998
207	1609	1608	4001	4000
208	1611	1610	4003	4002
209	1613	1612	4005	4004
210	1615	1614	4007	4006
211	1617	1616	4009	4008
212	1619	1618	4011	4010
213	1621	1620	4013	4012
214	1623	1622	4015	4014
215	1625	1624	4017	4016
216	1627	1626	4019	4018
217	1629	1628	4021	4020
218	1631	1630	4023	4022
219	1633	1632	4025	4024
220	1635	1634	4027	4026
221	1637	1636	4029	4028
222	1639	1638	4031	4030
223	1641	1640	4033	4032
224	1643	1642	4035	4034
225	1645	1644	4037	4036
226	1647	1646	4039	4038
227	1649	1648	4041	4040
228	1651	1650	. 4043	4042
229	1653	1652	4045	4044
230	1655	1654	4047	4046
231	1657	1656	4049	4048
232	1659	1658	4051	4050
233	1661	1660	4053	4052
234	1663	1662	4055	4054
235	1665	1664	4057	4056
236	1667	1666	4059	4058
237	1669	1668	4061	4060

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
238	1671	1670	4063	4062
239	1673	1672	4065	4064
240	1675	1674	4067	4066
241	1677	1676	4069	4068
242	1679	1678	4071	4070
243	1681	1680	4073	4072
244	1683	1682	4075	4074
245	1685	1684	4077	4076
246	1687	1686	4079	4078
247	1689	1688	4081	4080
248	1691	1690	4083	4082
249	1693	1692	4085	4084
250	1695	1694	4087	4086
251	1697	1696	4089	4088
252	1699	1698	4091	4090
253	1701	1700	4093	4092
254	1703	1702	4095	4094
255	1705	1704	4097	4096
256	1707	1706	4099	4098
257	1709	1708	4101	4100
258	1711	1710	4103	4102
259	1713	1712	4105	4104
260	1715	1714	4107	4106
261	1717	1716	4109	4108
262	1719	1718	4111	4110
263	1721	1720	4113	4112
264	1723	1722	4115	4114
265	1725	1724	4117	4116
266	1727	1726	4119	4118
267	1729	1728	4121	4120
268	1731	1730	4123	4122
269	1733	1732	4125	4124
270	1735	1734	4127	4126
271	1737	1736	4129	4128

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
272	1739	1738	4131	4130
273	1741	1740	4133	4132
274	1743	1742	4135	4134
275	1745	1744	4137	4136
276	1747	1746	4139	4138
277	1749	1748	4141	4140
278	1751	1750	4143	4142
279	1753	1752	4145	4144
280	1755	1754	4147	4146
281	1757	1756	4149	4148
282	1759	1758	4151	4150
283	1761	1760	4153	4152
284	1763	1762	4155	4154
285	1765	1764	4157	4156
286	1767	1766	4159	4158
287	1769	1768	4161	4160
288	1771	1770	4163	4162
289	1773	1772	4165	4164
290	1775	1774	4167	4166
291	1777	1776	4169	4168
292	1779	1778	4171	4170
293	1781	1780	4173	4172
294	1783	1782	4175	4174
295	1785	1784	4177	4176
296	1787	1786	4179	4178
297	1789	1788	4181	4180
298	1791	1790	4183	4182
299	1793	1792	4185	4184
300	1795	1794	4187	4186
301	1797	1796	4189	4188
302	1799	1798	4191	4190
303	1801	1800	4193	4192
304	1803	1802	4195	4194
305	1805	1804	4197	4196

ORFGenset .	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
306	1807	1806	4199	4198
307	1809	1808	4201	4200
308	1811	1810	4203	4202
309	1813	1812	4205	4204
310	1815	1814	4207	4206
311	1817	1816	4209	4208
312	1819	1818	4211	4210
313	1821	1820	4213	4212
314	1823	1822	4215	4214
315	1825	1824	4217	4216
316	1827	1826	4219	4218
317	1829	1828	4221	4220
318	1831	1830	4223	4222
319	1833	1832	4225	4224
320	1835	1834	4227	4226
321	1837	1836	4229	4228
322	1839	1838	4231	4230
323	1841	1840	4233	4232
324	1843	1842	4235	4234
325	1845	1844	4237	4236
326	1847	1846	4239	4238
327	1849	1848	4241	4240
328	1851	1850	4243	4242
329	1853	1852	4245	4244
330	1855	1854	4247	4246
331	1857	1856	4249	4248
332	1859	1858	4251	4250
333	1861	1860	4253	4252
334	1863	1862	4255	4254
335	1865	1864	4257	4256
336	1867	1866	4259	4258
337	1869	1868	4261	4260
338	1871	1870	4263	4262
339	1873	1872	4265	4264

ORFGenset	ORFoligosFd	ORFoligasFp	ORFoligosBd	ORFöligösBp
340	1875	1874	4267	4266
341	1877	1876	4269	4268
342	1879	1878	4271	4270
343	1881	1880	4273	4272
344	1883	1882	4275	4274
345	1885	1884	4277	4276
346	1887	1886	4279	4278
347	1889	1888	4281	4280
348	1891	1890	4283	4282
349	1893	1892	4285	4284
350	1895	1894	4287	4286
351	1897	1896	4289	4288
352	1899	1898	4291	4290
353	1901	1900	4293	4292
354	1903	1902	4295	4294
355	1905	1904	4297	4296
356	1907	1906	4299	4298
357	1909	1908	4301	4300
358	1911	1910	4303	4302
359	1913	1912	4305	4304
360	1915	1914	4307	4306
361	1917	1916	4309	4308
362	1919	1918	4311	4310
363	1921	1920	4313	4312
364	1923	1922	4315	4314
365	1925	1924	4317	4316
366	1927	1926	4319	4318
367	1929	1928	4321	4320
368	1931	1930	4323	4322
369	1933	1932	4325	4324
370	1935	1934	4327	4326
371	1937	1936	4329	4328
372	1939	1938	4331	4330
373	1941	1940	4333	4332

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
374	1943	1942	4335	4334
375	1945	1944	4337	4336
376	1947	1946	4339	4338
377	1949	1948	4341	4340
378	1951	1950	4343	4342
379	1953	1952	4345	4344
380	1955	1954	4347	4346
381	1957	1956	4349	4348
382	1959	1958	4351	4350
383	1961	1960	4353	4352
384	1963	1962	4355	4354
385	1965	1964	4357	4356
386	1967	1966	4359	4358
387	1969	1968	4361	4360
388	1971	1970	4363	4362
389	1973	1972	4365	4364
390	1975	1974	4367	4366
391	1977	1976	4369	4368
392	1979	1978	4371	4370
393	1981	1980	4373	4372
394	1983	1982	4375	4374
395	1985	1984	4377	4376
396	1987	1986	4379	4378
397	1989	1988	4381	4380
398	1991	1990	4383	4382
399	1993	1992	4385	4384
400	1995	1994	4387	4386
401	1997	1996	4389	4388
402	1999	1998	4391	4390
403	2001	2000	4393	4392
404	2003	2002	4395	4394
405	2005	2004	4397	4396
406	2007	2006	4399	4398
407	2009	2008	4401	4400

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
408	2011	2010	4403	4402
409	2013	2012	4405	4404
410	2015	2014	4407	4406
411	2017	2016	4409	4408
412	2019	2018	4411	4410
413	2021	2020	4413	4412
414	2023	2022	4415	4414
415	2025	2024	4417	4416
416	2027	2026	4419	4418
417	2029	2028	4421	4420
418	2031	2030	4423	4422
419	2033	2032	4425	4424
420	2035	2034	4427	4426
421	2037	2036	4429	4428
422	2039	2038	4431	4430
423	2041	2040	4433	4432
424	2043	2042	4435	4434
425	2045	2044	4437	4436
426	2047	2046	4439	4438
427	2049	2048	4441	4440
428	2051	2050	4443	4442
429	2053	2052	4445	4444
430	2055	2054	4447	4446
431	2057	2056	4449	4448
432	2059	2058	4451	4450
433	2061	2060	4453	4452
434	2063	2062	4455	4454
435	2065	2064	4457	4456
436	2067	2066	4459	4458
437	2069	2068	4461	4460
438	2071	2070	4463	4462
439	2073	2072	4465	4464
440	2075	2074	4467	4466
441	2077	2076	4469	4468

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
442	2079	2078	4471	4470
443	2081	2080	4473	4472
444	2083	2082	4475	4474
445	2085	2084	4477	4476
446	2087	2086	4479	4478
447	2089	2088	4481	4480
448	2091	2090	4483	4482
449	2093	2092	4485	4484
450	2095	2094	4487	4486
451	2097	2096	4489	4488
452	2099	2098	4491	4490
453	2101	2100	4493	4492
454	2103	2102	4495	4494
455	2105	2104	4497	4496
456	2107	2106	4499	4498
457	2109	2108	4501	4500
458	2111	2110	4503	4502
459	2113	2112	4505	4504
460	2115	2114	4507	4506
461	2117	2116	4509	4508
462	2119	2118	4511	4510
463	2121	2120	4513	4512
464	2123	2122	4515	4514
465	2125	2124	4517	4516
466	2127	2126	4519	4518
467	2129	2128	4521	4520
468	2131	2130	4523	4522
469	2133	2132	4525	4524
470	2135	2134	4527	4526
471	2137	2136	4529	4528
472	2139	2138	4531	4530
473	2141	2140	4533	4532
474	2143	2142	4535	4534
475	2145	2144	4537	4536

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligasBp
476	2147	2146	4539	4538
477	2149	2148	4541	4540
478	2151	2150	4543	4542
479	2153	2152	4545	4544
480	2155	2154	4547	4546
481	2157	2156	4549	4548
482	2159	2158	4551	4550
483	2161	2160	4553	4552
484	2163	2162	4555	4554
485	2165	2164	4557	4556
486	2167	2166	4559	4558
487	2169	2168	4561	4560
488	2171	2170	4563	4562
489	2173	2172	4565	4564
490	2175	2174	4567	4566
491	2177	2176	4569	4568
492	2179	2178	4571	4570
493	2181	2180	4573	4572
494	2183	2182	4575	4574
495	2185	2184	4577	4576
496	2187	2186	4579	4578
497	2189	2188	4581	4580
498	2191	2190	4583	4582
499	2193	2192	4585	4584
500	2195	2194	4587	4586
501	2197	2196	4589	4588
502	2199	2198	4591	4590
503	2201	2200	4593	4592
504	2203	2202	4595	4594
505	2205	2204	4597	4596
506	2207	2206	4599	4598
507	2209	2208	4601	4600
508	2211	2210	4603	4602
509	2213	2212	4605	4604

ORFGenset:	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
510	2215	2214	4607	4606
511	2217	2216	4609	4608
512	2219	2218	4611	4610
513	2221	2220	4613	4612
514	2223	2222	4615	4614
515	2225	2224	4617	4616
516	2227	2226	4619	4618
517	2229	2228	4621	4620
518	2231	2230	4623	4622
519	2233	2232	4625	4624
520	2235	2234	4627	4626
521	2237	2236	4629	4628
522	2239	2238	4631	4630
523	2241	2240	4633	4632
524	2243	2242	4635	4634
525	2245	2244	4637	4636
526	2247	2246	4639	4638
527	2249	2248	4641	4640
528	2251	2250	4643	4642
529	2253	2252	4645	4644
530	2255	2254	4647	4646
531	2257	2256	4649	4648
532	2259	2258	4651	4650
533	2261	2260	4653	4652
534	2263	2262	4655	4654
535	2265	2264	4657	4656
536	2267	2266	4659	4658
537	2269	2268	4661	4660
538	2271	2270	4663	4662
539	2273	2272	4665	4664
540	2275	2274	4667	4666
541	2277	2276	4669	4668
542	2279	2278	4671	4670
543	2281	2280	4673	4672

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
544	2283	2282	4675	4674
545	2285	2284	4677	4676
546	2287	2286	4679	4678
547	2289	2288	4681	4680
548	2291	2290	4683	4682
549	2293	2292	4685	4684
550	2295	2294	4687	4686
551	2297	2296	4689	4688 -
552	2299	2298	4691	4690
553	2301	2300	4693	4692
554	2303	2302	4695	4694
555	2305	2304	4697	4696
556	2307	2306	4699	4698
557	2309	2308	4701	4700
558	2311	2310	4703	4702
559	2313	2312	4705	4704
560	2315	2314	4707	4706
561	2317	2316	4709	4708
562	2319	2318	4711	4710
563	2321	2320	4713	4712
564	2323	2322	4715	4714
565	2325	2324	4717	4716
566	2327	2326	4719	4718
567	2329	2328	4721	4720
568	2331	2330	4723	4722
569	2333	2332	4725	4724
570	2335	2334	4727	4726
571	2337	2336	4729	4728
572	2339	2338	4731	4730
573	2341	2340	4733	4732
574	2343	2342	4735	4734
575	2345	2344	4737	4736
576	2347	2346	4739	4738
577	2349	2348	4741	4740

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
578	2351	2350	4743	4742
579	2353	2352	4745	4744
580	2355	2354	4747	4746
581	2357	2356	4749	4748
582	2359	2358	4751	4750
583	2361	2360	4753	4752
584	2363	2362	4755	4754
585	2365	2364	4757	4756
586	2367	2366	4759	4758
587	2369	2368	4761	4760
588	2371	2370	4763	4762
589	2373	2372	4765	4764
590	2375	2374	4767	4766
591	2377	2376	4769	4768
592	2379	2378	4771	4770
593	2381	2380	4773	4772
594	2383	2382	4775	4774
595	2385	2384	4777	4776
596	2387	2386	4779	4778
597	2389	2388	4781	4780
598	2391	2390	4783	4782
599	2393	2392	4785	4784
600	2395	2394	4787	4786
601	2397	2396	4789	4788
602	2399	2398	4791	4790
603	2401	2400	4793	4792
604	2403	2402	4795	4794
605	2405	2404	4797	4796
606	2407	2406	4799	4798
607	2409	2408	4801	4800
608	2411	2410	4803	4802
609	2413	2412	4805	4804
610	2415	2414	4807	4806
611	2417	2416	4809	4808

ORFGenset	ORFoligosFd	ORFöligösFp	ORFoligosBd	ORFoligosBp
612	2419	2418	4811	4810
613	2421	2420	4813	4812
614	2423	2422	4815	4814
615	2425	2424	4817	4816
616	2427	2426	4819	4818
617	2429	2428	4821	4820
618	2431	2430	4823	4822
619	2433	2432	4825	4824
620	2435	2434	4827	4826
621	2437	2436	4829	4828
622	2439	2438	4831	4830
623	2441	2440	4833	4832
624	2443	2442	4835	4834
625	2445	2444	4837	4836
626	2447	2446	4839	4838
627	2449	2448	4841	4840
628	2451	2450	4843	4842
629	2453	2452	4845	4844
630	2455	2454	4847	4846
631	2457	2456	4849	4848
632	2459	2458	4851	4850
633	2461	2460	4853	4852
634	2463	2462	4855	4854
635	2465	2464	4857	4856
636	2467	2466	4859	4858
637	2469	2468	4861	4860
638	2471	2470	4863	4862
639	2473	2472	4865	4864
640	2475	2474	4867	4866
641	2477	2476	4869	4868
642	2479	2478	4871	4870
643	2481	2480	4873	4872
644	2483	2482	4875	4874
645	2485	2484	4877	4876

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
646	2487	2486	4879	4878
647	2489	2488	4881	4880
648	2491	2490	4883	4882
649	2493	2492	4885	4884
650	2495	2494	4887	4886
651	2497	2496	4889	4888
652	2499	2498	4891	4890
653	2501	2500	4893	4892
654	2503	2502	4895	4894
655	2505	2504	4897	4896
656	2507	2506	4899	4898
657	2509	2508	4901	4900
658	2511	2510	4903	4902
659	2513	2512	4905	4904
660	2515	2514	4907	4906
661	2517	2516	4909	4908
662	2519	2518	4911	4910
663	2521	2520	4913	4912
664	2523	2522	4915	4914
665	2525	2524	4917	4916
666	2527	2526	4919	4918
667	2529	2528	4921	4920
668	2531	2530	4923	4922
669	2533	2532	4925	4924
670	2535	2534	4927	4926
671	2537	2536	4929	4928
672	2539	2538	4931	4930
673	2541	2540	4933	4932
674	2543	2542	4935	4934
675	2545	2544	4937	4936
676	2547	2546	4939	4938
677	2549	2548	4941	4940
678	2551	2550	4943	4942
679	2553	2552	4945	4944

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
680	2555	2554	4947	4946
681	2557	2556	4949	4948
682	2559	2558	4951	4950
683	2561	2560	4953	4952
684	2563	2562	4955	4954
685	2565	2564	4957	4956
686	2567	2566	4959	4958
687	2569	2568	4961	4960
688	2571	2570	4963	4962
689	2573	2572	4965	4964
690	2575	2574	4967	4966
691	2577	2576	4969	4968
692	2579	2578	4971	4970
693	2581	2580	4973	4972
694	2583	2582	4975	4974
695	2585	2584	4977	4976
696	2587	2586	4979	4978
697	2589	2588	4981	4980
698	2591	2590	4983	4982
699	2593	2592	4985	4984
700	2595	2594	4987	4986
701	2597	2596	4989	4988
702	2599	2598	4991	4990
703	2601	2600	4993	4992
704	2603	2602	4995	4994
705	2605	2604	4997	4996
706	2607	2606	4999	4998
707	2609	2608	5001	5000
708	2611	2610	5003	5002
709	2613	2612	5005	5004
710	2615	2614	5007	5006
711	2617	2616	5009	5008
712	2619	2618	5011	5010
713	2621	2620	5013	5012

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
714	2623	2622	5015	5014
715	2625	2624	5017	5016
716	2627	2626	5019	5018
717	2629	2628	5021	5020
718	2631	2630	5023	5022
719	2633	2632	5025	5024
720	2635	2634	5027	5026
721	2637	2636	5029	5028
722	2639	2638	5031	5030
723	2641	2640	5033	5032
724	2643	2642	5035	5034
725	2645	2644	5037	5036
726	2647	2646	5039	5038
727	2649	2648	5041	5040
728	2651	2650	5043	5042
729	2653	2652	5045	5044
730	2655	2654	5047	5046
731	2657	2656	5049	5048
732	2659	2658	5051	5050
733	2661	2660	5053	5052
734	2663	2662	5055	5054
735	2665	2664	5057	5056
736	2667	2666	5059	5058
737	2669	2668	5061	5060
738	2671	2670	5063	5062
739	2673	2672	5065	5064
740	2675	2674	5067	5066
741	2677	2676	5069	5068
742	2679	2678	5071	5070
743	2681	2680	5073	5072
744	2683	2682	5075	5074
745	2685	2684	5077	5076
746	2687	2686	5079	5078
747	2689	2688	5081	5080

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
748	2691	2690	5083	5082
749	2693	2692	5085	5084
750	2695	2694	5087	5086
751	2697	2696	5089	5088
752	2699	2698	5091	5090
753	2701	2700	5093	5092
754	2703	2702	5095	5094
755	2705	2704	5097	5096
756	2707	2706	5099	5098
757	2709	2708	5101	5100
758	2711	2710	5103	5102
759	2713	2712	5105	5104
760	2715	2714	5107	5106
761	2717	2716	5109	5108
762	2719	2718	5111	5110
763	2721	2720	5113	5112
764	2723	2722	5115	5114
765	2725	2724	5117	5116
766	2727	2726	5119	5118
767	2729	2728	5121	5120
768	2731	2730	5123	5122
769	2733	2732	5125	5124
770	2735	2734	5127	5126
771	2737	2736	5129	5128
772	2739	2738	5131	5130
773	2741	2740	5133	5132
774	2743	2742	5135	5134
775	2745	2744	5137	5136
776	2747	2746	5139	5138
777	2749	2748	5141	5140
778	2751	2750	5143	5142
779	2753	2752	5145	5144
780	2755	2754	5147	5146
781	2757	2756	5149	5148

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
782	2759	2758	5151	5150
783	2761	2760	5153	5152
784	2763	2762	5155	5154
785	2765	2764	5157	5156
786	2767	2766	5159	5158
787	2769	2768	5161	5160
788	2771	2770	5163	5162
789	2773	2772	5165	5164
790	2775	2774	5167	5166
791	2777	2776	5169	5168
792	2779	2778	5171	5170
793	2781	2780	5173	5172
794	2783	2782	5175	5174
795	2785	2784	5177	5176
796	2787	2786	5179	5178
797	2789	2788	5181	5180
798	2791	2790	5183	5182
799	2793	2792	5185	5184
800	2795	2794	5187	5186
801	2797	2796	5189	5188
802	2799	2798	5191	5190
803	2801	2800	5193	5192
804	2803	2802	5195	5194
805	2805	2804	5197	5196
806	2807	2806	5199	5198
807	2809	2808	5201	5200
808	2811	2810	5203	5202
809	2813	2812	5205	5204
810	2815	2814	5207	5206
811	2817	2816	5209	5208
812	2819	2818	5211	5210
813	2821	2820	5213	5212
814	2823	2822	5215	5214
815	2825	2824	5217	5216

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
816	2827	2826	5219	5218
817	2829	2828	5221	5220
818	2831	2830	5223	5222
819	2833	2832	5225	5224
820	2835	2834	5227	5226
821	2837	2836	5229	5228
822	2839	2838	5231	5230
823	2841	2840	5233	5232
824	2843	2842	5235	5234
825	2845	2844	5237	5236
826	2847	2846	5239	5238
827	2849	2848	5241	5240
828	2851	2850	5243	5242
829	2853	2852	5245	5244
830	2855	2854	5247	5246
831	2857	2856	5249	5248
832	2859	2858	5251	5250
833	2861	2860	5253	5252
834	2863	2862	5255	5254
835	2865	2864	5257	5256
836	2867	2866	5259	5258
837	2869	2868	5261	5260
838	2871	2870	5263	5262
839	2873	2872	5265	5264
840	2875	2874	5267	5266
841	2877	2876	5269	5268
842	2879	2878	5271	5270
843	2881	2880	5273	5272
844	2883	2882	5275	5274
845	2885	2884	5277	5276
846	2887	2886	5279	5278
847	2889	2888	5281	5280
848	2891	2890	5283	5282
849	2893	2892	5285	5284

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
850	2895	2894	5287	5286
851	2897	2896	5289	5288
852	2899	2898	5291	5290
853	2901	2900	5293	5292
854	2903	2902	5295	5294
855	2905	2904	5297	5296
856	2907	2906	5299	5298
857	2909	2908	5301	5300
858	2911	2910	5303	5302
859	2913	2912	5305	5304
860	2915	2914	5307	5306
861	2917	2916	5309	5308
862	2919	2918	5311	5310
863	2921	2920	5313	5312
864	2923	2922	5315	5314
865	2925	2924	5317	5316
866	2927	2926	5319	5318
867	2929	2928	5321	5320
868	2931	2930	5323	5322
869	2933	2932	5325	5324
870	2935	2934	5327	5326
871	2937	2936	5329	5328
872	2939	2938	5331	5330
873	2941	2940	5333	5332
874	2943	2942	5335	5334
875	2945	2944	5337	5336
876	2947	2946	5339	5338
877	2949	2948	5341	5340
878	2951	2950	5343	5342
879	2953	2952	5345	5344
880	2955	2954	5347	5346
881	2957	2956	5349	5348
882	2959	2958	5351	5350
883	2961	2960	5353	5352

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
884	2963	2962	5355	5354
885	2965	2964	5357	5356
886	2967	2966	5359	5358
887	2969	2968	5361	5360
888	2971	2970	5363	5362
889	2973	2972	5365	5364
890	2975	2974	5367	5366
891	2977	2976	5369	5368
892	2979	2978	5371	5370
893	2981	2980	5373	5372
894	2983	2982	5375	5374
895	2985	2984	5377	5376
896	2987	2986	5379	5378
897	2989	2988	5381	5380
898	2991	2990	5383	5382
899	2993	2992	5385	5384
900	2995	2994	5387	5386
901	2997	2996	5389	5388
902	2999	2998	5391	5390
903	3001	3000	5393	5392
904	3003	3002	5395	5394
905	3005	3004	5397	5396
906	3007	3006	5399	5398
907	3009	3008	5401	5400
908	3011	3010	5403	5402
909	3013	3012	5405	5404
910	3015	3014	5407	5406
911	3017	3016	5409	5408
912	3019	3018	5411	5410
913	3021	3020	5413	5412
914	3023	3022	5415	5414
915	3025	3024	5417	5416
916	3027	3026	5419	5418
917	3029	3028	5421	5420

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
918	3031	3030	5423	5422
919	3033	3032	5425	5424
920	3035	3034	5427	5426
921	3037	3036	5429	5428
922	3039	3038	5431	5430
923	3041	3040	5433	5432
924	3043	3042	5435	5434
925	3045	3044	5437	5436
926	3047	3046	5439	5438
927	3049	3048	5441	5440
928	3051	3050	5443	5442
929	3053	3052	5445	5444
930	3055	3054	5447	5446
931	3057	3056	5449	5448
932	3059	3058	5451	5450
933	3061	3060	5453	5452
934	3063	3062	5455	5454
935	3065	3064	5457	5456
936	3067	3066	5459	5458
937	3069	3068	5461	5460
938	3071	3070	5463	5462
939	3073	3072	5465	5464
940	3075	3074	5467	5466
941	3077	3076	5469	5468
942	3079	3078	5471	5470
943	3081	3080	5473	5472
944	3083	3082	5475	5474
945	3085	3084	5477	5476
946	3087	3086	5479	5478
947	3089	3088	5481	5480
948	3091	3090	5483	5482
949	3093	3092	5485	5484
950	3095	3094	5487	5486
951	3097	3096	5489	5488

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
952	3099	3098	5491	5490
953	3101	3100	5493	5492
954	3103	3102	5495	5494
955	3105	3104	5497	5496
956	3107	3106	5499	5498
957	3109	3108	5501	5500
958	3111	3110	5503	5502
959	3113	3112	5505	5504
960	3115	3114	5507	5506
961	3117	3116	5509	5508
962	3119	3118	5511	5510
963	3121	3120	5513	5512
964	3123	3122	5515	5514
965	3125	3124	5517	5516
966	3127	3126	5519	5518
967	3129	3128	5521	5520
968	3131	3130	5523	5522
969	3133	3132	5525	5524
970	3135	3134	5527	5526
971	3137	3136	5529	5528
972	3139	3138	5531	5530
973	3141	3140	5533	5532
974	3143	3142	5535	5534
975	3145	3144	5537	5536
976	3147	3146	5539	5538
977	3149	3148	5541	5540
978	3151	3150	5543	5542
979	3153	3152	5545	5544
980	3155	3154	5547	5546
981	3157	3156	5549	5548
982	3159	3158	5551	5550
983	3161	3160	5553	5552
984	3163	3162	5555	5554
985	3165	3164	5557	5556

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
986	3167	3166	5559	5558
987	3169	3168	5561	5560
988	3171	3170	5563	5562
989	3173	3172	5565	5564
990	3175	3174	5567	5566
991	3177	3176	5569	5568
992	3179	3178	5571	5570
993	3181	3180	5573	5572
994	3183	3182	5575	5574
995	3185	3184	5577	5576
996	3187	3186	5579	5578
997	3189	3188	5581	5580
998	3191	3190	5583	5582
999	3193	3192	5585	5584
1000	3195	3194	5587	5586
1001	3197	3196	5589	5588
1002	3199	3198	5591	5590
1003	3201	3200	5593	5592
1004	3203	3202	5595	5594
1005	3205	3204	5597	5596
1006	3207	3206	5599	5598
1007	3209	3208	5601	5600
1008	3211	3210	5603	5602
1009	3213	3212	5605	5604
1010	3215	3214	5607	5606
1011	3217	3216	5609	5608
1012	3219	3218	5611	5610
1013	3221	3220	5613	5612
1014	3223	3222	5615	5614
1015	3225	3224	5617	5616
1016	3227	3226	5619	5618
1017	3229	3228	5621	5620
1018	3231	3230	5623	5622
1019	3233	3232	5625	5624

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1020	3235	3234	5627	5626
1021	3237	3236	5629	5628
1022	3239	3238	5631	5630
1023	3241	3240	5633	5632
1024	3243	3242	5635	5634
1025	3245	3244	5637	5636
1026	3247	3246	5639	5638
1027	3249	3248	5641	5640
1028	3251	3250	5643	5642
1029	3253	3252	5645	5644
1030	3255	3254	5647	5646
1031	3257	325ú	5649	5648
1032	3259	3258	5651	5650
1033	3261	3260	5653	5652
1034	3263	3262	5655	5654
1035	3265	3264	5657	5656
1036	3267	3266	5659	5658
1037	3269	3268	5661	5660
1038	3271	3270	5663	5662
1039	3273	3272	5665	5664
1040	3275	3274	5667	5666
1041	3277	3276	5669	5668
1042	3279	3278	5671	5670
1043	3281	3280	5673	5672
1044	3283	3282	5675	5674
1045	3285	3284	5677	5676
1046	3287	3286	5679	5678
1047	3289	3288	5681	5680
1048	3291	3290	5683	5682
1049	3293	3292	5685	5684
1050	3295	3294	5687	5686
1051	3297	3296	5689	5688
1052	3299	3298	5691	5690
1053	3301	3300	5693	5692

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1054	3303	3302	5695	5694
1055	3305	3304	5697	5696
1056	3307	3306	5699	5698
1057	3309	3308	5701	5700
1058	3311	3310	5703	5702
1059	3313	3312	5705	5704
1060	3315	3314	5707	5706
1061	3317	3316	5709	5708
1062	3319	3318	5711	5710
1063	3321	3320	5713	5712
1064	3323	3322	5715	5714
1065	3325	3324	5717	5716
1066	3327	3326	5719	5718
1067	3329	3328	5721	5720
1068	3331	3330	5723	5722
1069	3333	3332	5725	5724
1070	3335	3334	5727	5726
1071	3337	3336	5729	5728
1072	3339	3338	5731	5730
1073	3341	3340	5733	5732
1074	3343	3342	5735	5734
1075	3345	3344	5737	5736
1076	3347	3346	5739	5738
1077	3349	3348	5741	5740
1078	3351	3350	5743	5742
1079	3353	3352	5745	5744
1080	3355	3354	5747	5746
1081	3357	3356	5749	5748
1082	3359	3358	5751	5750
1083	3361	3360	5753	5752
1084	3363	3362	5755	5754
1085	3365	3364	5757	5756
1086	3367	3366	5759	5758
1087	3369	3368	5761	5760

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1088	3371	3370	5763	5762
1089	3373	3372	5765	5764
1090	3375	3374	5767	5766
1091	3377	3376	5769	5768
1092	3379	3378	5771	5770
1093	3381	3380	5773	5772
1094	3383	3382	5775	5774
1095	3385	3384	5777	5776
1096	3387	3386	5779	5778
1097	3389	3388	5781	5780
1098	3391	3390	5783	5782
1099	3393	3392	5785	5784
1100	3395	3394	5787	5786
1101	3397	3396	5789	5788
1102	3399	3398	5791	5790
1103	3401	3400	5793	5792
1104	3403	3402	5795	5794
1105	3405	3404	5797	5796
1106	3407	3406	5799	5798
1107	3409	3408	5801	5800
1108	3411	3410	5803	5802
1109	3413	3412	5805	5804
1110	3415	3414	5807	5806
1111	3417	3416	5809	5808
1112	3419	3418	5811	5810
1113	3421	3420	5813	5812
1114	3423	3422	5815	5814
1115	3425	3424	5817	5816
1116	3427	3426	5819	5818
1117	3429	3428	5821	5820
1118	3431	3430	5823	5822
1119	3433	3432	5825	5824
1120	3435	3434	5827	5826
1121	3437	3436	5829	5828

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1122	3439	3438	5831	5830
1123	3441	3440	5833	5832
1124	3443	3442	5835	5834
1125	3445	3444	5837	5836
1126	3447	3446	5839	5838
1127	3449	3448	5841	5840
1128	3451	3450	5843	5842
1129	3453	3452	5845	5844
1130	3455	3454	5847	5846
1131	3457	3456	5849	5848
1132	3459	3458	5851	5850
1133	3461	3460	5853	5852
1134	3463	3462	5855	5854
1135	3465	3464	5857	5856
1136	3467	3466	5859	5858
1137	3469	3468	5861	5860
1138	3471	3470	5863	5862
1139	3473	3472	5865	5864
1140	3475	3474	5867	5866
1141	3477	3476	5869	5868
1142	3479	3478	5871	5870
1143	3481	3480	5873	5872
1144	3483	3482	5875	5874
1145	3485	3484	5877	5876
1146	3487	3486	5879	5878
1147	3489	3488	5881	5880
1148	3491	3490	5883	5882
1149	3493	3492	5885	5884
1150	3495	3494	5887	5886
1151	3497	3496	5889	5888
1152	3499	3498	5891	5890
1153	3501	3500	5893	5892
1154	3503	3502	5895	5894
1155	3505	3504	5897	5896

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1156	3507	3506	5899	5898
1157	3509	3508	5901	5900
1158	3511	3510	5903	5902
1159	3513	3512	5905	5904
1160	3515	3514	5907	5906
1161	3517	3516	5909	5908
1162	3519	3518	5911	5910
1163	3521	3520	5913	5912
1164	3523	3522	5915	5914
1165	3525	3524	5917	5916
1166	3527	3526	5919	5918
1167	3529	3528	5921	5920
1168	3531	3530	5923	5922
1169	3533	3532	5925	5924
1170	3535	3534	5927	5926
1171	3537	3536	5929	5928
1172	3539	3538	5931	5930
1173	3541	3540	5933	5932
1174	3543	3542	5935	5934
1175	3545	3544	5937	5936
1176	3547	3546	5939	5938
1177	3549	3548	5941	5940
1178	3551	3550	5943	5942
1179	3553	3552	5945	5944
1180	3555	3554	5947	5946
1181	3557	3556	5949	5948
1182	3559	3558	5951	5950
1183	3561	3560	5953	5952
1184	3563	3562	5955	5954
1185	3565	3564	5957	5956
1186	3567	3566	5959	5958
1187	3569	3568	5961	5960
1188	3571	3570	5963	5962
1189	3573	3572	5965	5964

ORFGenset	ORFoligosFd	ORFoligosFp	ORFoligosBd	ORFoligosBp
1190	3575	3574	5967	5966
1191	3577	3576	5969	5968
1192	3579	3578	5971	5970
1193	3581	3580	5973	5972
1194	3583	3582	5975	5974
1195	3585	3584	5977	5976
1196	3587	3586	5979	,5978
1197	3589	3588	5981	5980

#### TABLE 5

SEQ ID	Or	. position
1198	F	1038449
1199	F	1036517
1200	F	250
1201	F	1036965
1202	F	3011
1203	F	1123
1204	F	4907
1205	F	2996
1206	F	6379
1207	F	4483
1208	F	7837
1209	F	5961
1210	F	8351
1211	F	6467
1212	F	8705
1213	F	6834
1214	F	9598
1215	F	7709
1216	F	10134
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SEQ ID	Or.	position
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4734       B       564319         4735       B       566220         4736       B       566220         4737       B       566078         4738       B       566078         4739       B       567243         4740       B       569143         4740       B       568192         4741       B       570619         4742       B       572532         4744       B       572241         4745       B       574208         4746       B       572994         4747       B       574916         4748       B       573744         4749       B       575640         4750       B       573679         4751       B       575571         4752       B       574398         4753       B       576288         4754       B       576633         4755       B       576922         4758       B       575482	4732	В	563153
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3203       F       968888         3204       F       972169         3205       F       970269         3206       F       973487         3207       F       971616         3208       F       974339         3209       F       974988         3210       F       974988         3211       F       976035         3212       F       976367         3213       F       974114         3214       F       976367         3215       F       974411         3216       F       976665         3217       F       974730         3218       F       977439         3219       F       975500         3220       F       977698         3221       F       975799         3222       F       978389         3223       F       976478         3224       F       978665         3225       F       976760         3226       F       979134         3227       F       977592         3230       F       980915 <t< td=""><td>3201</td><td>F</td><td>967320</td></t<>	3201	F	967320
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3206       F       973487         3207       F       971616         3208       F       974339         3209       F       972408         3210       F       974988         3211       F       976035         3212       F       976035         3213       F       974114         3214       F       976367         3215       F       974411         3216       F       976665         3217       F       974730         3218       F       977439         3219       F       975500         3220       F       977698         3221       F       975799         3222       F       978389         3223       F       976478         3224       F       978665         3225       F       976760         3226       F       979134         3227       F       977592         3230       F       980915         3231       F       979026	3204	F	972169
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3212 F 976035 3213 F 974114 3214 F 976367 3215 F 974411 3216 F 976665 3217 F 974730 3218 F 977439 3219 F 975500 3220 F 977698 3221 F 975799 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3210	F	974988
3213 F 974114 3214 F 976367 3215 F 974411 3216 F 976665 3217 F 974730 3218 F 977439 3219 F 975500 3220 F 977698 3221 F 975799 3222 F 978389 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3211	F	973035 .
3214 F 976367  3215 F 974411  3216 F 976665  3217 F 974730  3218 F 977439  3219 F 975500  3220 F 977698  3221 F 975799  3222 F 978389  3223 F 976478  3224 F 978665  3225 F 976760  3226 F 979134  3227 F 977270  3228 F 979473  3229 F 977592  3230 F 980915  3231 F 979026	3212	F	976035
3215 F 974411 3216 F 976665 3217 F 974730 3218 F 977439 3219 F 975500 3220 F 977698 3221 F 975799 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3213	F	974114
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3217 F 974730 3218 F 977439 3219 F 975500 3220 F 977698 3221 F 975799 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3215	F	974411
3218 F 977439 3219 F 975500 3220 F 977698 3221 F 975799 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3216	F	976665
3219 F 975500  3220 F 977698  3221 F 975799  3222 F 978389  3223 F 976478  3224 F 978665  3225 F 976760  3226 F 979134  3227 F 977270  3228 F 979473  3229 F 977592  3230 F 980915  3231 F 979026	3217	F	974730
3220 F 977698  3221 F 975799  3222 F 978389  3223 F 976478  3224 F 978665  3225 F 976760  3226 F 979134  3227 F 977270  3228 F 979473  3229 F 977592  3230 F 980915  3231 F 979026	3218	F	977439
3221 F 975799 3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3219	F	975500
3222 F 978389 3223 F 976478 3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3220	F	977698
3223 F 976478  3224 F 978665  3225 F 976760  3226 F 979134  3227 F 977270  3228 F 979473  3229 F 977592  3230 F 980915  3231 F 979026	3221	F	975799
3224 F 978665 3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3222	F	978389
3225 F 976760 3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3223	F	976478
3226 F 979134 3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3224	F	978665
3227 F 977270 3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026		F	976760
3228 F 979473 3229 F 977592 3230 F 980915 3231 F 979026	3226	F	979134
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3230 F 980915 3231 F 979026		F	979473
3231 F 979026		F	977592
	3230	F	980915
3232 F 982043	3231	F	979026
	3232	Ŧ	982043

SEQ ID	Or.	position
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4797	В	594885
4798	В	593367
4799	В	595321
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4803	В	597826
4804	В	596354
4805	В	598255
4806	В	597147
4807	В	598998
4808	В	597960
4809	В	599851
4810	В	601068
4811	В	602929
4812	В	602096
4813	В	603996
4814	В	603761
4815	В	605643
4816	В	604014
4817	В	605920
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SEQ ID	Or.	position
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1640	F	221314
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1643	F	220338
1644	F	223186
1645	F	221278
1646	F	223994
1647	F	222146
1648	F	224908
1649	F	223014
1650	F	225051
1651	F	223131
1652	F	225510
1653	F	223615
1654	F	226550
1655	F	224609
1656	F	226928
1657	F	225029
1658	F	227528
1659	F	225631
1660	F	228388
1661	F	226475
1662	F	229930
1663	F	228032
1664	F	231129
1665	F	229205
1666	F	232785
1667	F	230915
1668	F	233561
1669	F	231664
1670	F	234013
1671	F	232149

SEQ ID	Or.	position
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3235	F	981736
3236	F	984149
3237	F	982271
3238	F	985180
3239	F	983280
3240	F	985815
3241	F	983882
3242	F	986458
3243	F	984547
3244	F	987340
3245	F	985462
3246	F	987686
3247	F	985815
3248	F	988559
3249	F	986670
3250	F	989455
3251	F	987558
3252	F	993122
3253	F	991289
3254	F	993186
3255	F	991297
3256	F	993650
3257	F	991727
3258	F	994007
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3260	F	995743
3261	F	993915
3262	F	996663
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3265	F	996755
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SEQ ID	Or.	position
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4831	В	613998
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4833	В	615091
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4835	В	615761
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4837	В	616094
4838	В	615464
4839	В	617391
4840	В	615913
4841	В	617803
4842	В	617932
4843	В	619837
4844	В	618598
4845	В	620532
4846	В	619591
4847	В	621465
4848	В	620636
4849	В	622536
4850	В	620532
4851	В	622381
4852	В	621404
4853	В	623307
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4855	В	623834
4856	В	622666
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4859	В	625605
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SEQ ID	Or.	position
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1674	F	236015
1675	F	234123
1676	F	237945
1677	F	236045
1678	F	238482
1679	F	236599
1680	F	240094
1681	F	238190
1682	F	241713
1683	F	239820
1684	F	242569
1685	F	240653
1686	F	244253
1687	F	242360
1688	F	245693
1689	F	243796
1690	F	246762
1691	F	244825
1692	F	247498
1693	F	245575
1694	F	248343
1695	F	246444
1696	F	249500
1697	F	247625
1698	F	250315
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1702	F	251847
1703	F	249939
1704	F	254897
1705	F	252955

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SEQ ID	Or.	position	
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3268	F	1000967	
3269	F	999092	
3270	F	1001173	
3271	F	999246	
3272	F	1001604	
3273	F	999645	
3274	F	1004159	
3275	F	1002326	
3276	F	1004763	
3277	F	1002871	
3278	F	1005160	
3279	F	1003235	
3280	F	1007181	
3281	F	1005250	
3282	F	1007561	
3283	F	1005665	
3284	F	1008855	
3285	F	1007002	
3286	F	1010205	
3287	F	1008342	
3288	F	1011716	
3289	F	1009823	
3290	F	1011812	
3291	F	1009914	
3292	F	1012372	
3293	F	1010385	
3294	F	1012567	
3295	F	1010624	
3296	F	1013237	
3297	F	1011337	
3298	F	1013690	
3299	F	1011856	
3300	F	1014301	

SEQ ID	Or.	. position
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4864	В	626885
4865	В	628790
4866	В	627128
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4868	В	628073
4869	В	629983
4870	В	628359
4871	В	630267
4872	В	628976
4873	В	63085C
4874	В	630023
4875	В	631988
4876	В	630642
4877	В	632526
4878	В	631205
4879	В	633081
4880	В	632046
4881	В	633969
4882	В	638311
4883	В	640204
4884	В	640448
4885	В	642328
4886	В	643695
4887	В	645639
4888	В	640847
4889	В	642769
4890	В	644745
4891	В	646615
4892	В	645686
4893	В	647558
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SEQ ID	Or.	position
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1707	F	254643
1708	F	257692
1709	F	255790
1710	F	258561
1711	F	256651
1712	F	258927
1713	F	257036
1714	F	261368
1715	F	259469
1716	F	263887
1717	F	262000
1718	F	264503
1719	F	262599
1720	F	265364
1721	F	263512
1722	F	266202
1723	F	264277
1724	F	266709
1725	F	264801
1726	F	267847
1727	F	265947
1728	F	267980
1729	F	266077
1730	F	268271
1731	F	266341
1732	F	269840
1733	F	267913
1734	F	270961
1735	F	269072
1736	F	271883
1737	F	270080
1738	F	272642
1739	F	270748

SEQ ID	10-	
	Or	
3301	F	1012396
3302	F	1014926
3303	F	1013010
3304	F	1015664
3305	F	1013820
3306	F	1017026
3307	F	1015099
3308	F	1017674
3309	F	1015786
3310	F	1018353
3311	F	1016460
3312	F	1019602
3313	F	1017674
3314	F	1019876
3315	F	1017948
3316	F	1020853
3317	F	1018956
3318	F	1021878
3319	F	1019972
3320	F	1023054
3321	F	1021186
3322	F	1023415
3323	F	1021579
3324	F	1023748
3325	F	1021850
3326	F	1024485
3327	F	1022574
3328	F	1024744
3329	F	1022836
3330	F	1025618
3331	F	1023720
3332	F	1026323
3333	F	1024403
3334	F	1027710

SEQ ID	Or	
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4899	В	651829
4900	В	650580
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4902	В	651942
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4906	В	653132
4907	В	655028
4908	В	653827
4909	В	655713
4910	В	662071
4911	В	664023
4912	В	662543
4913	В	664403
4914	В	663295
4915	В	665205
4916	В	663972
4917	В	665850
4918	В	664432
4919	В	666332
4920	В	665860
4921	В	667789
4922	В	666312
4923	В	668233
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4927	В	670238
4928	В	668605
4929	В	670495

SEQ ID	Or.	position
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1742	F	274562
1743	F	272702
1744	F	275882
1745	F	273984
1746	F	278004
1747	F	276149
1748	F	278747
1749	F	276893
1750	F	279521
1751	F	277632
1752	F	281076
1753	F	279118
1754	F	281551
1755	F	279668
1756	F	282573
1757	F	280663
1758	F	284229
1759	F	282316
1760	F	284598
1761	F	282655
1762	F	285418
1763	F	283518
1764	F	286104
1765	F	284229
1766	F	286456
1767	F	284531
1768	F	287865
1769	F	286008
1770	F	289163
1771	F	287384
1772	F	290609
1773	F	288709

SEQ ID	Or.	position
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3336	F	1030272
3337	F	1028389
3338	F	1031486
3339	F	1029602
3340	F	1033215
3341	F	1031334
3342	F	1035425
3343	F	1033555
3344	F	1035956
3345	F	1034055
3346	F	1036748
3347	F	1034844
3348	F	16372
3349	F	14463
3350	F	31184
3351	F	29287
3352	F	56283
3353	F	54383
3354	F	56384
3355	F	54538
3356	F	64528
3357	F	62600
3358	F	72965
3359	F	71054
3360	F	78245
3361	F	76347
3362	F	79133
3363	Ŧ	77291
3364	F	81740
3365	F	79840
3366	F	86772
3367	F	84880
3368	F	109188

SEQ ID	Or.	p sition
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4935	В	672109
4936	В	671000
4937	В	672900
4938	В	671470
4939	В	673412
4940	В	672685
4941	В	674567
4942	В	673461
4943	В	675365
4944	В	674786
4945	В	676682
4946	В	675456
4947	В	677375
4948	В	676683
4949	В	678594
4950	В	677334
4951	В	679183
4952	В	678726
4953	В	680596
4954	В	679729
4955	В	681628
4956	В	680747
4957	В	682668
4958	В	681500
4959	В	683406
4960	В	682779
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4962		683320
4963	В	685249
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SEQ ID         Or.         position           1774         F         291264           1775         F         289389           1776         F         292107           1777         F         290166           1778         F         293099           1779         F         291211           1780         F         294791           1781         F         292883           1782         F         295464           1783         F         293573           1784         F         296018           1785         F         294095           1786         F         297572           1787         F         295664           1788         F         296716           1790         F         300305           1791         F         298407           1792         F         301852           1793         F         299946
1775         F         289389           1776         F         292107           1777         F         290166           1778         F         293099           1779         F         291211           1780         F         294791           1781         F         292883           1782         F         295464           1783         F         293573           1784         F         296018           1785         F         294095           1786         F         297572           1787         F         295664           1788         F         298686           1789         F         296716           1790         F         300305           1791         F         298407           1792         F         301852
1776       F       292107         1777       F       290166         1778       F       293099         1779       F       291211         1780       F       294791         1781       F       292883         1782       F       295464         1783       F       293573         1784       F       296018         1785       F       294095         1786       F       297572         1787       F       295664         1788       F       298686         1789       F       296716         1790       F       300305         1791       F       298407         1792       F       301852
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1778         F         293099           1779         F         291211           1780         F         294791           1781         F         292883           1782         F         295464           1783         F         293573           1784         F         296018           1785         F         294095           1786         F         297572           1787         F         295664           1788         F         298686           1789         F         296716           1790         F         300305           1791         F         298407           1792         F         301852
1779       F       291211         1780       F       294791         1781       F       292883         1782       F       295464         1783       F       293573         1784       F       296018         1785       F       294095         1786       F       297572         1787       F       295664         1788       F       298686         1789       F       296716         1790       F       300305         1791       F       298407         1792       F       301852
1780       F       294791         1781       F       292883         1782       F       295464         1783       F       293573         1784       F       296018         1785       F       294095         1786       F       297572         1787       F       295664         1788       F       298686         1789       F       296716         1790       F       300305         1791       F       298407         1792       F       301852
1781       F       292883         1782       F       295464         1783       F       293573         1784       F       296018         1785       F       294095         1786       F       297572         1787       F       295664         1788       F       298686         1789       F       296716         1790       F       300305         1791       F       298407         1792       F       301852
1782 F 295464  1783 F 293573  1784 F 296018  1785 F 294095  1786 F 297572  1787 F 295664  1788 F 298686  1789 F 296716  1790 F 300305  1791 F 298407  1792 F 301852
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SEQ ID	Or.	position
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3371	F	109188
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3373	F	109597
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3380	F	124892
3381	F	123004
3382	F	131327
3383	F	129485
3384	F	143944
3385	F	142043
3386	F	150138
3387	F	148247
3388	F	163715
3389	F	161804
3390	F	165186
3391	F	163274
3392	F	168143
3393	F	166302
3394	F	170287
3395	F	168387
3396	F	176838
3397	F	174996
3398	F	187776
3399	F	185900
3400	F	188083
3401	F	186208
3402	F	190117

4964       B       684716         4965       B       686585         4966       B       685010         4967       B       686897         4968       B       686423         4969       B       688323         4970       B       687426         4971       B       688619         4972       B       688619         4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4987       B       696078         4989       B       697093         4990       B       697486         4991       B       698313 <t< th=""><th>SEQ ID</th><th>Or.</th><th>position</th></t<>	SEQ ID	Or.	position
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4968       B       686423         4969       B       688323         4970       B       687426         4971       B       689324         4972       B       688619         4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692088         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4994       B       698646 <t< td=""><td>4966</td><td>В</td><td>685010</td></t<>	4966	В	685010
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4970       B       687426         4971       B       689324         4972       B       688619         4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697486         4991       B       698313         4992       B       698313         4993       B       698646         4995       B       700515         4996       B       700337 <td>4968</td> <td>В</td> <td>686423</td>	4968	В	686423
4971       B       689324         4972       B       688619         4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4993       B       698646         4995       B       700515         4996       B       700337	4969	В	688323
4972       B       688619         4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4993       B       698646         4995       B       700515         4996       B       700337	4970	В	687426
4973       B       690482         4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4971	В	689324
4974       B       688653         4975       B       690563         4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4972	В	688619
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4976       B       689836         4977       B       691775         4978       B       690186         4979       B       692088         4980       B       692715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698313         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4974	В	688653
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4978       B       690186         4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698428         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4976	В	689836
4979       B       692088         4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       699428         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4977	В	691775
4980       B       690715         4981       B       692616         4982       B       690937         4983       B       692837         4984       B       692091         4985       B       693991         4986       B       694171         4987       B       696078         4988       B       695197         4989       B       697093         4990       B       697486         4991       B       698428         4992       B       698313         4993       B       700238         4994       B       698646         4995       B       700515         4996       B       700337	4978	В	690186
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4997 B 702249	4996	В	700337
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SEQ ID	Or.	position
1808	F	311753
1809	F	309790
1810	F	313188
1811	F	311292
1812	F	314121
1813	F	312194
1814	F	314489
1815	F	312539
1816	F	315431
1817	F	313526
1818	F	316309
1819	F	314380
1820	F	317102
1821	F	315214
1822	F	317271
1823	F	315343
1824	F	317380
1825	F	315480
1826	F	318256
1827	F	316352
1828	F	319047
1829	F	317181
1830	F	320325
1831	F	318338
1832	F	321228
1833	F	319366
1834	F	321676
1835	F	319782
1836	F	322066
1837	F	320097
1838	F	322910
1839	F	320982
1840	F	324744
1841	F	322849
		· · · · · ·

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3410	F	236682
3411	F	234794
3412	F	249227
3413	F	247310
3414	F	252939
3415	F	251036
3416	F	253406
3417	F	251562
3418	F	271365
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3420	F	275390
3421	F	273489
3422	F	277681
3423	F	275765
3424	F	282260
3425	F	280357
3426	F	292925
3427	F	291054
3428	F	302910
3429	F	301032
3430	F	308746
3431	F	306806
3432	F	311994
3433	F	310073
3434	F	312375
3435	F	310483
3436	F	312531

SEQ ID	Or.	position
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4999	В	703015
5000	В	702385
5001	В	704285
5002	В	703636
5003	В	705561
5004	В	705271
5005	В	707136
5006	В	705875
5007	В	707725
5008	В	706444
5009	В	708279
5010	В	706741
5011	В	708673
5012	В	708324
5013	В	710226
5014	В	708673
5015	В	710518
5016	В	708876
5017	В	710791
5018	В	710498
5019	В	712447
5020	В	711435
5021	В	713354
5022	В	712993
5023	В	714887
5024	В	713686
5025	В	715574
5026	В	714474
5027	В	716354
5028	В	714867
5029	В	716760
5030		716047
5031	В	717877

SEQ ID	Or.	position
1842	F	325392
1843	F	323445
1844	F	326217
1845	F	324331
1846	F	327038
1847	F	325162
1848	F	327957
1849	F	326079
1850	F	328458
1851	F	326612
1852	F	329032
1853	F	327173
1854	F	329329
1855	F	327489
1856	F	330446
1857	F	328551
1858	F	330915
1859	F	329032
1860	F	331410
1861	F	329602
1862	F	332534
1863	F	330626
1864	F	332782
1865	F	330879
1866	F	333587
1867	F	331632
1868	F	333870
1869	F	331962
1870	F	334510
1871	F	332594
1872	F	334958
1873	F	333049
1874	F	334958
1875	F	333049

SEQ ID	Or.	position
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3438	F	319923
3439	F	318009
3440	F	339991
3441	F	338104
3442	F	352535
3443	F	350653
3444	F	373218
3445	F	371320
3446	F	376994
3447	F	375085
3448	F	378954
3449	F	377011
3450	F	394604
3451	F	392704
3452	F	400915
3453	F	398972
3454	F	409744
3455	F	407904
3456	F	411155
3457	F	409253
3458	F	414197
3459	F	412281
3460	F	422638
3461	F	420770
3462	F	427595
3463	F	425701
3464	F	428453
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3466	F	442272
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3470	F	442939
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SEQ ID	Or.	position
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5036	В	718624
5037	В	720503
5038	В	719083
5039	В	720983
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5041	В	722004
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5043	В	722393
5044	В	720753
5045	В	722653
5046	В	721798
5047	В	723724
5048	В	722631
5049	В	724493
5050	В	723468
5051	В	725376
5052	В	724852
5053	В	726743
5054	В	726005
5055	В	727903
5056	В	726779
5057	В	728691
5058	В	727058
5059	В	728947
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5063	В	730116
5064	В	729048
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SEQ ID	Or.	position
1876	F	335655
1877	F	333766
1878	F	336117
1879	F	334219
1880	F	337108
1881	F	335210
1882	F	340251
1883	F	338372
1884	F	341538
1885	F	339662
1886	F	341953
1887	F	339995
1888	F	342348
1889	F	340450
1890	F	343112
1891	F	341242
1892	F	343736
1893	F	341811
1894	F	344117
1895	F	342207
1896	F	344940
1897	F	343000
1898	F	345837
1899	F	343958
1900	F	346872
1901	F	344994
1902	F	347910
1903	F	345971
1904	F	350124
1905	F	348298
1906	F	351095
1907	F	349167
1908	F	351996
1909	F	350122

3471       F       441047         3472       F       445572         3473       F       443707         3474       F       467757         3475       F       465801         3476       F       471583         3477       F       469712         3478       F       487813         3479       F       485913         3480       F       496852         3481       F       499979         3482       F       499979         3483       F       498074         3484       F       506798         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       564318         3495       F       564318         3496       F       564838         3498       F       570033 <t< th=""><th>SEQ ID</th><th>Or.</th><th>position</th></t<>	SEQ ID	Or.	position
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3474       F       467757         3475       F       465801         3476       F       471583         3477       F       469712         3478       F       487813         3479       F       485913         3480       F       496852         3481       F       494952         3482       F       499979         3483       F       498074         3484       F       506798         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       564318         3494       F       564318         3495       F       564838         3498       F       570033         3498       F       570844         3500       F       575571         3503       F       573671 <td>3472</td> <td>F</td> <td>445572</td>	3472	F	445572
3475       F       465801         3476       F       471583         3477       F       469712         3478       F       487813         3479       F       485913         3480       F       496852         3481       F       499979         3482       F       499979         3483       F       498074         3484       F       508715         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       554706         3494       F       564318         3495       F       566692         3497       F       564838         3498       F       570033         3499       F       568150         3500       F       575571         3503       F       573671 <td>3473</td> <td>F</td> <td>443707</td>	3473	F	443707
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3478       F       487813         3479       F       485913         3480       F       496852         3481       F       494952         3482       F       499979         3483       F       498074         3484       F       508715         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       556575         3494       F       564318         3495       F       566692         3497       F       564838         3498       F       570033         3499       F       568150         3500       F       570844         3501       F       575571         3503       F       573671	3476	F	471583
3479       F       485913         3480       F       496852         3481       F       494952         3482       F       499979         3483       F       498074         3484       F       508715         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       556575         3494       F       564318         3495       F       566692         3497       F       564838         3498       F       570033         3499       F       568915         3500       F       575571         3503       F       573671	3477	F	469712
3480       F       496852         3481       F       494952         3482       F       499979         3483       F       498074         3484       F       508715         3485       F       506798         3486       F       510584         3487       F       508632         3488       F       526255         3489       F       524350         3490       F       531098         3491       F       529150         3492       F       556575         3493       F       554706         3494       F       564318         3495       F       566692         3497       F       564838         3498       F       570033         3499       F       568150         3500       F       570844         3501       F       568915         3502       F       573671	3478	F	487813
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3491 F 529150  3492 F 556575  3493 F 554706  3494 F 564318  3495 F 562390  3496 F 566692  3497 F 564838  3498 F 570033  3499 F 568150  3500 F 570844  3501 F 568915  3502 F 575571  3503 F 573671	3489	F	524350
3492 F 556575  3493 F 554706  3494 F 564318  3495 F 562390  3496 F 566692  3497 F 564838  3498 F 570033  3499 F 568150  3500 F 570844  3501 F 568915  3502 F 575571  3503 F 573671	3490	F	531098
3493 F 554706  3494 F 564318  3495 F 562390  3496 F 566692  3497 F 564838  3498 F 570033  3499 F 568150  3500 F 570844  3501 F 568915  3502 F 575571  3503 F 573671	3491	F	529150
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	3502	F	575571
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SEQ ID	Or.	position
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1911	F	351186
1912	F	353413
1913	F	351481
1914	F	353908
1915	F	351996
1916	F	354723
1917	F	352799
1918	F	356466
1919	F	354569
1920	F	357107
1921	F	355178
1922	F	357767
1923	F	355878
1924	F	360528
1925	F	358628
1926	F	360877
1927	F	358974
1928	F	361573
1929	F	359692
1930	F	362584
1931	F	360681
1932	F	363835
1933	F	361966
1934	F	364960
1935	F	363021
1936	F	365240
1937	F	363360
1938	F	367060
1939	F	365115
1940	F	368383
1941	F	366505
1942	F	368862
1943	F	366963

	Or.	position
3505	F	588196
3506	F	597631
3507	F	595698
3508	F	606387
3509	F	604507
3510	F	607566
3511	F	605637
3512	F	609842
3513	F	607958
3514	F	632472
3515	F	630572
3516	F	636994
3517	F	635071
3518	F	649681
3519	F	647800
3520	F	652059
3521	F	650101
3522	F	654522
3523	F	652562
3524	F	660587
3525	F	658691
3526	F	676785
3527	F	674938
3528	F	679031
3529	1	677133
3530	F	731967
3531	F	730091
3532	F	741797
3533	F	739935
3534	F	758555
3535	F	756641
3536	F	760010
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SEQ ID	Or.	position
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5102	В	748519
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5105	В	751108
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5111	В	752151
5112	В	752527
5113	В	754427
5114	В	753169
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5116	В	755004
5117	В	756843
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5119	В	758991
5120	В	758532
5121	В	760452
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SEQ ID	Or.	position
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1945	F	368631
1946	F	370974
1947	F	369076
1948	F	372891
1949	F	370980
1950	F	373395
1951	F	371495
1952	F	374005
1953	F	372033
1954	F	374474
1955	F	372572
1956	F.	376509
1957	F	374624
1958	F	377630
1959	F	375708
1960	F	378384
1961	F	376507
1962	F	378798
1963	F	376871
1964	F	379413
1965	F	377501
1966	F	379890
1967	F	377989
1968	F	381241
1969	F	379348
1970	F	382485
1971	F	380579
1972	F	383395
1973	F	381536
1974	F	383730
1975	F	381782
1976	F	384948
1977	F	383057

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SEQ ID	Or.	ļ <sup>*</sup>
3539	F	768751
3540	F	771896
3541	F	769996
3542	F	787857
3543	F	785958
3544	F	815714
3545	F	813840
3546	F	846380
3547	F	844470
3548	F	867576
3549	F	865673
3550	F	875167
3551	F	873254
3552	F	876214
3553	F	874314
3554	F	884093
3555	F	882162
3556	F	891248
3557	F	889348
3558	F	900125
3559	F	898298
3560	F	902048
3561	F	900125
3562	F	907563
3563	F	905656
3564	F	912076
3565	F	910133
3566	F	935157
3567	F	933211
3568	F	946473
3569	F	944568
3570	F	952562
3571	F	950664
3572	F	965649
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SEQ ID	Or.	position
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5135	B	765107
5136	В	764690
5137	В	766595
5138	В	765107
5139	В	766977
5140	В	766327
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5142	В	766932
5143	В	768851
5144	В	768314
5145	В	770221
5146	В	769045
5147	В	770945
5148	В	770315
5149	В	772234
5150	В	770705
5151	В	772598
5152	В	770882
5153	В	772781
5154	В	771156
5155	В	773044
5156	В	772234
5157	В	774148
5158	В	773611
5159	В	775511
5160	В	774513
5161	В	776404
5162	В	776333
5163	В	778191
5164		777926
5165		779832
5166		777455
5167	В	779380

SEQ ID	Or.	position
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1979	F	383532
1980	F	385908
1981	F	384008
1982	F	386643
1983	F	384750
1984	F	387099
1985	F	385204
1986	F	387581
1987	F	385677
1988	F	388009
1989	F	386062
1990	F	388927
1991	F	387033
1992	F	389726
1993	F	387821
1994	F	391295
1995	F	389365
1996	F	392171
1997	F	390291
1998	F	393930
1999	F	392014
2000	F	395085
2001	F	393185
2002	F	395827
2003	F	393940
2004	F	396274
2005	F	394423
2006	F	397156
2007	F	395216
2008	F	398641
2009	F	396790
2010	F	399550
2011	F	397659

SEQ ID	Or.	position
3573	F	963730
3574	F	968519
3575	F	966614
3576	F	970497
3577	F	968601
3578	F	971879
3579	F	970043
3580	F	972888
3581	F	970962
3582	F	998162
3583	F	996241
3584	F	1003657
3585	F	1001756
3586	F	1009313
3587	F	1007413
3588	F	1028954
3589	F	1027039
3590	В	730
3591	В	2645
3592	В	3521
3593	В	5431
3594	В	5295
3595	В	7188
3596	В	6740
3597	В	8652
3598	В	8240
3599	В	10138
3600	В	8959
3601	В	10816
3602	В	9285
3603	В	11160
3604	В	9689
3605	В	11591
3606	В	10679

SEQ ID	Or	position
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5169	В	781342
5170	В	781774
5171	В	783686
5172	В	782667
5173	В	784562
5174	В	785748
5175	В	787658
5176	В	786222
5177	В	788126
5178	В	786803
5179	В	788703
5180	В	787998
5181	В	789876
5182	В	788279
5183	В	790255
5184	В	790369
5185	В	792247
5186	В	790862
5187	В	792787
5188	В	792247
5189	В	794137
5190	В	793352
5191	В	795215
5192	В	794276
5193	В	796196
5194	В	795215
5195	В	797077
5196	В	795667
5197	В	797571
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SEQ ID	Or.	position
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2013	F	397915
2014	F	401527
2015	F	399623
2016	F	401907
2017	F	399960
2018	F	403017
2019	F	401131
2020	F	403017
2021	F	401131
2022	F	404910
2023	F	403010
2024	F	405728
2025	F	403836
2026	F	406837
2027	F	404932
2028	F	410291
2029	F	408347
2030	F	411488
2031	F	409518
2032	F	412379
2033	F	410487
2034	F	413164
2035	F	411263
2036	F	413606
2037	F	411626
2038	F	413721
2039	F	411859
2040	F	414921
2041	F	413049
2042	F	416517
2043	F	414606
2044	F	417445
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	SEQ ID	Or	. position
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	3608	В	11515
Ì	3609	В	13376
I	3610	В	12136
	3611	В	14038
	3612	В	13565
	3613	В	15450
	3614	В	14667
	3615	В	16535
	3616	В	15254
	3617	В	17183
	3618	В	16189
ſ	3619	В	18097
	3620	В	16790
ſ	3621	В	18621
ľ	3622	В	16710
ſ	3623	В	18587
	3624	В	18458
	3625	В	20318
Γ	3626	В	20768
	3627	В	22647
Γ	3628	В	22438
	3629	В	24335
	3630	В	22892
Г	3631	В	24752
Γ	3632	В	23046
	3633	В	24924
Γ	3634	В	23442
Γ	3635	В	25313
	3636	В	23623
	3637	В	25528
	3638	В	24822
	3639	В	26708
	3640	В	26618

SEQ ID	Or.	position
5202	В	797585
5203	В	799553
5204	В	798102
5205	В	799976
5206	В	798306
5207	В	800214
5208	В	798662
5209	В	800546
5210	В	799168
5211	В	801074
5212	В	800159
5213	В	802054
5214	В	801153
5215	В	803031
5216	В	802363
5217	В	804257
5218	В	802893
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	2066	F	429303
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Ţ	2068	F	429609
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3656	В	34121
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SEQ ID	Or	. position
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3680       B       45451         3681       B       47375         3682       B       46308         3683       B       48208         3684       B       46560         3685       B       48485         3686       B       47115         3687       B       49053         3688       B       48208         3689       B       50118         3690       B       48806         3691       B       50708         3692       B       50333         3693       B       52220         3694       B       50960         3695       B       52890         3696       B       52660         3697       B       54606         3698       B       53737         3699       B       55645         3700       B       54793         3701       B       56691         3704       B       56691         3706       B       58566         3706       B       56317         3707       B       58226	3678	В	44022
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5279         B         831395           5280         B         831192           5281         B         833082           5282         B         832005           5283         B         833870           5284         B         832373           5285         B         834280           5286         B         832964           5287         B         834864           5288         B         833672           5289         B         835604           5290         B         834022           5291         B         835945           5292         B         834907           5293         B         836803           5294         B         835683           5295         B         837593           5296         B         836018           5297         B         837902           5298         B         836714           5299         B         838602           5300         B         837484           5301         B         839424           5302         B         838950	5277	В	830176
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2158	F	476279
2159	F	474390
2160	F	478446
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2169	F	479377
2170	F	481635
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2188	F	492475
2189	F	490618
2190	F	493591
2191	F	491719
2192	F	494297
2193	F	492436
2194	F	494530
2195	F	492679
2196	F	494637
2197	F	492753
2198	F	495467
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2200	F	496076
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2202	F	497468
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2205	F	496160
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2208	F	500571
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SEQ ID	Or.	position
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3790	В	101639
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3792	В	102457
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3797	В	106464
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3800	В	106127
3801	В	108014
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537	9	В	880034
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538	1	В	880292
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538	3	В	881230
538	4	В	879393
538	5	В	881281
538	6	В	880089
538	7	В	881994
538	8	В	881108
538	9	В	883062
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2227	F	507356
2228	F	510759
2229	F	508918
2230	F	511268
2231	F	509359
2232	F	512124
2233	F	510202
2234	F	512836
2235	F	510926
2236	F	514569
2237	F	512663
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2239	F	512874
2240	F	515334
2241	F	513415
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2243	F	513670
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SEQ ID	Or	position
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5449       B       913064         5450       B       912492         5451       B       914381         5452       B       912852         5453       B       914763         5454       B       913438         5455       B       915340         5456       B       913901         5457       B       915814         5458       B       914730         5459       B       916694         5460       B       915356         5461       B       917256         5462       B       915717         5463       B       917608         5464       B       916854         5465       B       918761         5466       B       918762         5467       B       918712         5468       B       917848         5469       B       919756         5470       B       918588         5471       B       920424         5472       B       918875	5447	В	912689
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5451       B       914381         5452       B       912852         5453       B       914763         5454       B       913438         5455       B       915340         5456       B       915901         5457       B       915814         5458       B       914730         5459       B       916694         5460       B       915356         5461       B       917256         5462       B       915717         5463       B       917608         5464       B       916854         5465       B       918761         5466       B       916762         5467       B       918712         5468       B       917848         5469       B       919756         5470       B       918588         5471       B       920424         5472       B       918875	5449	В	913064
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5467     B     918712       5468     B     917848       5469     B     919756       5470     B     918588       5471     B     920424       5472     B     918875	5465	В	918761
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5469     B     919756       5470     B     918588       5471     B     920424       5472     B     918875	5467	В	918712
5470 B 918588 5471 B 920424 5472 B 918875	5468	В	917848
5471 B 920424 5472 B 918875	5469	В	919756
5472 B 918875	5470	В	918588
	5471	В	920424
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2284         F         532100           2285         F         530224           2286         F         533025           2287         F         531125           2288         F         533710           2289         F         531825           2290         F         534658           2291         F         532758           2292         F         535737           2293         F         53828           2294         F         539456           2295         F         537568           2296         F         540290           2297         F         538375           2298         F         540672           2299         F         538777           2300         F         541573           2301         F         543922           2303         F         541102           2304         F         543922           2305         F         542057           2306         F         544268           2307         F         542354           2308         F         54691           2310	SEQ ID	Or.	position
2286         F         533025           2287         F         531125           2288         F         533710           2289         F         531825           2290         F         534658           2291         F         532758           2292         F         535737           2293         F         539456           2294         F         539456           2295         F         537568           2296         F         540290           2297         F         538375           2298         F         540672           2299         F         538777           2300         F         541573           2301         F         542999           2302         F         542999           2303         F         541102           2304         F         542057           2305         F         542057           2306         F         544268           2307         F         542354           2308         F         544691           2310         F         546323           2311 <td>2284</td> <td>F</td> <td>532100</td>	2284	F	532100
2287         F         531125           2288         F         533710           2289         F         531825           2290         F         534658           2291         F         532758           2292         F         535737           2293         F         533828           2294         F         539456           2295         F         537568           2296         F         540290           2297         F         538375           2298         F         540672           2299         F         538777           2300         F         541573           2301         F         539706           2302         F         542999           2303         F         541102           2304         F         542057           2305         F         542057           2306         F         544268           2307         F         542354           2308         F         54691           2309         F         546323           2311         F         546467           2313 <td>2285</td> <td>F</td> <td>530224</td>	2285	F	530224
2288       F       533710         2289       F       531825         2290       F       534658         2291       F       532758         2292       F       535737         2293       F       533828         2294       F       539456         2295       F       537568         2296       F       540290         2297       F       538375         2298       F       540672         2299       F       538777         2300       F       541573         2301       F       539706         2302       F       542999         2303       F       541102         2304       F       543922         2305       F       542057         2306       F       544268         2307       F       542354         2308       F       544691         2309       F       546323         2310       F       546467         2313       F       546467         2313       F       545357         2316       F       547658 <td>2286</td> <td>F</td> <td>533025</td>	2286	F	533025
2289       F       531825         2290       F       534658         2291       F       532758         2292       F       535737         2293       F       535737         2294       F       539456         2295       F       537568         2296       F       540290         2297       F       538375         2298       F       540672         2299       F       538777         2300       F       541573         2301       F       539706         2302       F       542999         2303       F       541102         2304       F       543922         2305       F       542057         2306       F       544268         2307       F       542354         2308       F       544691         2309       F       546323         2311       F       546467         2313       F       54650         2314       F       547257         2315       F       547658	2287	F	531125
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2295         F         537568           2296         F         540290           2297         F         538375           2298         F         540672           2299         F         538777           2300         F         541573           2301         F         539706           2302         F         542999           2303         F         541102           2304         F         543922           2305         F         542057           2306         F         544268           2307         F         542354           2308         F         544691           2309         F         542785           2310         F         546323           2311         F         546467           2313         F         54650           2314         F         547658	2293	F	533828
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2297         F         538375           2298         F         540672           2299         F         538777           2300         F         541573           2301         F         539706           2302         F         542999           2303         F         541102           2304         F         543922           2305         F         542057           2306         F         544268           2307         F         542354           2308         F         544691           2309         F         542785           2310         F         546323           2311         F         546467           2313         F         544650           2314         F         547257           2315         F         545357           2316         F         547658	2295	F	537568
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5614       B       979204         5615       B       981104         5616       B       979554         5617       B       981465         5618       B       981423         5619       B       983319         5620       B       980363         5621       B       982289         5622       B       982361         5623       B       984236         5624       B       983818         5625       B       985718         5626       B       984720         5627       B       986608         5628       B       987553         5630       B       986323         5631       B       986925         5632       B       986925         5633       B       988925         5634       B       98749         5635       B       989749         5636       B       988474         5639       B       990252         5638       B       988474         5639       B       989437         5640       B       989437 <tr< td=""><td>5612</td><td>В</td><td>978655</td></tr<>	5612	В	978655
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4232	В	325882
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5920       B       679748         5921       B       681674         5922       B       732909         5923       B       734756         5924       B       742639         5924       B       742639         5925       B       744503         5926       B       759613         5927       B       761510         5928       B       760782         5929       B       762671         5930       B       771617         5931       B       772628         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       877779         5942       B       877737         5943       B       877737         5945       B       886680 <t< td=""><td>5918</td><td>В</td><td>677837</td></t<>	5918	В	677837
5921       B       681674         5922       B       732909         5923       B       734756         5924       B       742639         5925       B       744503         5926       B       759613         5927       B       761510         5928       B       760782         5929       B       762671         5930       B       771617         5931       B       772628         5932       B       774528         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5939       B       847145         5939       B       849042         5940       B       868276         5941       B       875887         5942       B       877137         5943       B       877137         5945       B       886680         5947       B       886680         5947       B       886680 <t< td=""><td>5919</td><td>В</td><td>679716</td></t<>	5919	В	679716
5922       B       732909         5923       B       734756         5924       B       742639         5925       B       744503         5926       B       759613         5927       B       761510         5928       B       760782         5929       B       762671         5930       B       771617         5931       B       772628         5932       B       774528         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       877137         5943       B       877137         5945       B       886680         5947       B       886680         5948       B       892172	5920	В	679748
5923       B       734756         5924       B       742639         5925       B       744503         5926       B       759613         5927       B       761510         5928       B       760782         5929       B       762671         5930       B       771617         5931       B       773519         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       877779         5943       B       877137         5945       B       886680         5947       B       886680         5947       B       886680         5948       B       892172	5921	В	681674
5924       B       742639         5925       B       744503         5926       B       759613         5927       B       761510         5928       B       760782         5929       B       762671         5930       B       771617         5931       B       772628         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       877779         5943       B       877137         5945       B       884780         5947       B       886680         5947       B       886680         5948       B       892172	5922	В	732909
5925         B         744503           5926         B         759613           5927         B         761510           5928         B         760782           5929         B         762671           5930         B         771617           5931         B         773519           5932         B         772628           5933         B         774528           5934         B         788703           5935         B         790577           5936         B         816591           5937         B         818443           5938         B         847145           5939         B         849042           5940         B         868276           5941         B         870177           5942         B         875887           5943         B         877137           5945         B         879035           5946         B         884780           5947         B         886680           5948         B         892172	5923	В	734756
5926         B         759613           5927         B         761510           5928         B         760782           5929         B         762671           5930         B         771617           5931         B         773519           5932         B         772628           5933         B         774528           5934         B         788703           5935         B         790577           5936         B         816591           5937         B         818443           5938         B         847145           5939         B         849042           5940         B         868276           5941         B         870177           5942         B         875887           5943         B         877137           5944         B         877137           5945         B         884780           5947         B         886680           5948         B         892172	5924	В	742639
5927         B         761510           5928         B         760782           5929         B         762671           5930         B         771617           5931         B         773519           5932         B         772628           5933         B         774528           5934         B         788703           5935         B         790577           5936         B         816591           5937         B         818443           5938         B         847145           5939         B         849042           5940         B         868276           5941         B         870177           5942         B         877779           5943         B         877137           5945         B         879035           5946         B         884780           5947         B         886680           5948         B         892172	5925	В	744503
5928         B         760782           5929         B         762671           5930         B         771617           5931         B         773519           5932         B         772628           5933         B         774528           5934         B         788703           5935         B         790577           5936         B         816591           5937         B         818443           5938         B         847145           5939         B         849042           5940         B         868276           5941         B         870177           5942         B         875887           5943         B         877779           5944         B         877137           5945         B         884780           5947         B         886680           5948         B         892172	5926	В	759613
5929       B       762671         5930       B       771617         5931       B       773519         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       884780         5947       B       886680         5948       B       892172	5927	В	761510
5930       B       771617         5931       B       773519         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877137         5944       B       877137         5945       B       884780         5947       B       886680         5948       B       892172	5928	В	
5931       B       773519         5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       884780         5947       B       886680         5948       B       892172	5929	В	762671
5932       B       772628         5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       884780         5947       B       886680         5948       B       892172		В	771617
5933       B       774528         5934       B       788703         5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       884780         5947       B       886680         5948       B       892172	5931	В	773519
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5935       B       790577         5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       879035         5946       B       884780         5947       B       886680         5948       B       892172	5933	В	774528
5936       B       816591         5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       879035         5946       B       884780         5947       B       886680         5948       B       892172	5934	В	788703
5937       B       818443         5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       879035         5946       B       884780         5947       B       886680         5948       B       892172	5935	В	790577
5938       B       847145         5939       B       849042         5940       B       868276         5941       B       870177         5942       B       875887         5943       B       877779         5944       B       877137         5945       B       879035         5946       B       884780         5947       B       886680         5948       B       892172	5936	В	816591
5939     B     849042       5940     B     868276       5941     B     870177       5942     B     875887       5943     B     877779       5944     B     877137       5945     B     879035       5946     B     884780       5947     B     886680       5948     B     892172		В	818443
5940     B     868276       5941     B     870177       5942     B     875887       5943     B     877779       5944     B     877137       5945     B     879035       5946     B     884780       5947     B     886680       5948     B     892172	5938	В	847145
5941     B     870177       5942     B     875887       5943     B     877779       5944     B     877137       5945     B     879035       5946     B     884780       5947     B     886680       5948     B     892172	5939	В	849042
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5944 B 877137 5945 B 879035 5946 B 884780 5947 B 886680 5948 B 892172	5942	В	
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5946 B 884780 5947 B 886680 5948 B 892172	_	В	877137
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	5947	В	
5949 B 894073		В	
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2773	F	774514
2774	F	776672
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2776	F	777446
2777	F	775596
2778	F	779102
2779	F	777192
2780	F	781078
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2782	F	782192
2783	F	780236
2784	F	785250
2785	F	783413
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2792	F	787693

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4363	В	385837
4364	В	384167
4365	В	386065
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4367	В	387365
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4372	В	386903
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4374	В	387595
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5975	В	1000774
5976	В	1004572
5977	В	1006449
5978	В	1010507
5979	В	1012353
5980	В	1029707
5981	В	1031628

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## WHAT IS CLAIMED IS:

- 1. An isolated polynucleotide having a nucleotide sequence of a Chlamydia trachomatis genome, comprising
  - (a) the nucleotide sequence of SEQ ID No. 1;
  - (b) the nucleotide sequence contained within the *Chlamydia trachomatis* genomic DNA in ECACC Deposit No. 98112618;
  - (c) the nucleotide sequence contained in a clone insert in ECACC Deposit No. 98112617;
  - (d) a nucleotide sequence exhibiting at least 99.9% identity with the sequence of SEQ ID No. 1; or
  - (e) a nucleotide sequence exhibiting at least 80% homology to SEQ ID No. 1.
- 2. An isolated polynucleotide which hybridizes to SEQ ID No. 1 or to the Chlamydia trachomatis genomic DNA contained in ECACC Deposit No. 98112618 or to a clone insert in ECACC Deposit No. 98112617 under conditions of high stringency.
- 3. An isolated polynucleotide which hybridizes to SEQ ID No. 1 or to the Chlamydia trachomatis genomic DNA contained in ECACC Deposit No. 98112618 under conditions of intermediate stringency.
- 4. An isolated polynucleotide having a nucleotide sequence of an open reading frame (ORF) of a *Chlamydia trachomatis* genome, comprising:
  - (a) a nucleotide sequence chosen from one of ORF2 to ORF 1197;
  - (b) a nucleotide sequence exhibiting at least 99.9% identity with one of ORF2 to ORF 1197; or
  - (c) a nucleotide sequence exhibiting at least 80% homology to one of ORF2 to ORF 1197.
- 5. An isolated polynucleotide which hybridizes to one of ORF2 to ORF 1197 under conditions of high stringency.
- 6. An isolated polynucleotide which hybridizes to one of ORF2 to ORF 1197 under conditions of intermediate stringency.
- 7. The polynucleotide of Claim 2, 3, 4, 5, or 6 which encodes the following polypeptides or fragments thereof:

- (a) a Chlamydia trachomatis transmembrane polypeptide having between 1 and 3 transmembrane domains;
- (b) a *Chlamydia trachomatis* transmembrane polypeptide having between 4 and 6 transmembrane domains;
- (c) a Chlamydia trachomatis transmembrane polypeptide having at least 7 transmembrane domains;
- a Chlamydia trachomatis polypeptide involved in intermediate metabolism of sugars and/or cofactors;
- (e) a Chlamydia trachomatis polypeptide involved in intermediate metabolism of nucleotides or nucleic acids;
- (f) a *Chlamydia trachomatis* polypeptide involved in metabolism of amino acids or polypeptides;
- (g) a Chlamydia trachomatis polypeptide having involved in metabolism of fatty acids;
- a Chlamydia trachomatis polypeptide involved in the synthesis of the cell wall;
- (i) a *Chlamydia trachomatis* polypeptide involved in transcription, translation, and/or maturation process;
- (j) a Chlamydia trachomatis transport polypeptide;
- (k) a Chlamydia trachomatis polypeptide involved in the virulence process;
- (l) a *Chlamydia trachomatis* polypeptide involved in the secretory system and/or which is secreted;
- (m) a Chlamydia trachomatis polypeptide of the cellular envelope or outer cellular envelope of Chlamydia trachomatis.
- (n) a Chlamydia trachomatis surface exposed polypeptide;
- (o) a Chlamydia trachomatis lipoprotein;
- (p) a *Chlamydia trachomatis* polypeptide involved in lipopolysaccharide biosynthesis;
- (q) a Chlamydia trachomatis KDO-related polypeptide;
- (r) a Chlamydia trachomatis phosphomannomutase-related polypeptide;
- (s) a Chlamydia trachomatis phosphoglucomutase-related polypeptide;
- (t) a Chlamydia trachomatis lipid A component-related polypeptide;
- (u) a Chlamydia trachomatis polypeptide that contains an RGD sequence;
- (v) a Chlamydia trachomatis Type III secreted polypeptide;
- (w) a Chlamydia trachomatis cell wall anchored surface polypeptide; or
- (x) a Chlamydia trachomatis polypeptide that is not found in Chlamydia trachomatis.

- 8. A polynucleotide encoding a fusion protein, comprising one of ORF2 to ORF 1197 of Claim 4, 5, or 6 ligated in frame to a polynucleotide encoding a heterologous polypeptide.
- 9. A recombinant vector that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
  - 10. A recombinant vector that contains the polynucleotide of Claim 8.
- 11. A recombinant vector that contains the polynucleotide of Claim 4, 5 or 6, operatively associated with a regulatory sequence that controls gene expression.
- 12. A recombinant vector that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression.
- 13. A genetically engineered host cell that contains the polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
  - 14. A genetically engineered host cell that contains the polynucleotide of Claim 8.
- 15. A genetically engineered host cell that contains the polynucleotide of Claim 4, 5 or 6 operatively associated with a regulatory sequence that controls gene expression in the host cell.
- 16. A genetically engineered host cell that contains the polynucleotide of Claim 8 operatively associated with a regulatory sequence that controls gene expression in the host cell.
  - 17. A method for producing a polypeptide, comprising:
  - (a) culturing the genetically engineered host cell of Claim 15 under conditions suitable to produce the polypeptide encoded by the polynucleotide; and
  - (b) recovering the polypeptide from the culture.
  - 18. A method for producing a fusion protein, comprising:
  - (a) culturing the genetically engineered host cell of Claim 16 under conditions suitable to produce the fusion protein encoded by the polynucleotide; and
  - (b) recovering the fusion protein from the culture.
  - 19. A polypeptide encoded by the polynucleotide of Claim 4, 5 or 6.

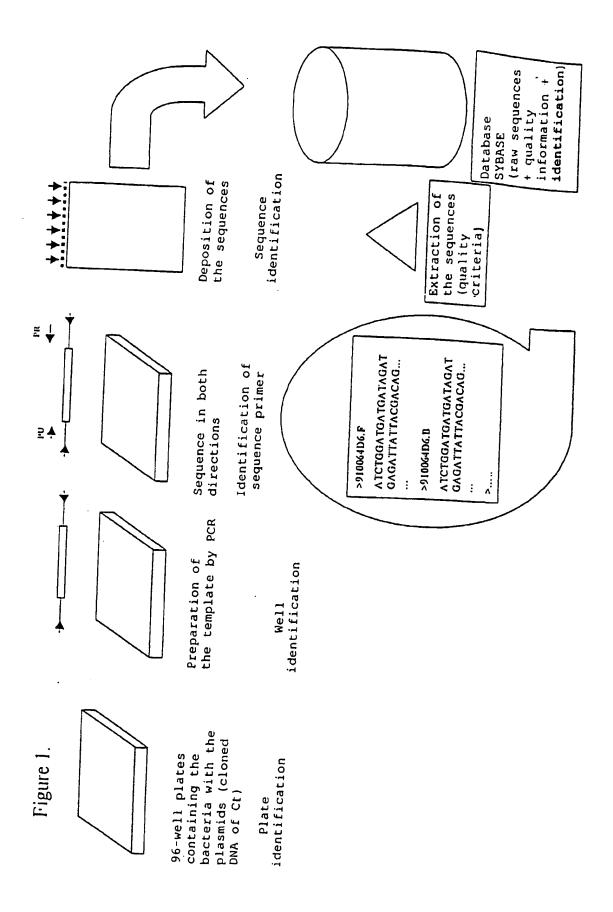
- 20. The polypeptide of Claim 19 which immunoreacts with seropositive serum of an individual infected with *Chlamydia trachomatis*.
- 21. The polypeptide of Claim 19 which comprises the following polypeptides or fragments thereof:
  - (a) a *Chlamydia trachomatis* transmembrane polypeptide having between 1 and 3 transmembrane domains;
  - (b) a *Chlamydia trachomatis* transmembrane polypeptide having between 4 and 6 transmembrane domains;
  - (c) a *Chlamydia trachomatis* transmembrane polypeptide having at least 7 transmembrane domains;
  - (d) a Chlamydia trachomatis polypeptide involved in intermediate metabolism of sugars and/or cofactors;
  - (e) a *Chlamydia trachomatis* polypeptide involved in intermediate metabolism of nucleotides or nucleic acids;
  - (f) a *Chlamydia trachomatis* polypeptide involved in metabolism of amino acids or polypeptides;
  - (g) a Chlamydia trachomatis polypeptide involved in metabolism of fatty acids;
  - a Chlamydia trachomatis polypeptide involved in the synthesis of the cell wall;
  - a Chlamydia trachomatis polypeptide involved in transcription, translation, and/or maturation process;
  - (j) a Chlamydia trachomatis transport polypeptide;
  - (k) a Chlamydia trachomatis polypeptide involved in the virulence process;
  - (l) a *Chlamydia trachomatis* polypeptide involved in the secretory system and/or which is secreted;
  - (m) a Chlamydia trachomatis polypeptide of the cellular envelope or outer cellular envelope of Chlamydia trachomatis.
  - (n) a Chlamydia trachomatis surface exposed polypeptide;
  - (o) a Chlamydia trachomatis lipoprotein;
  - a Chlamydia trachomatis polypeptide involved in lipopolysaccharide biosynthesis;
  - (q) a Chlamydia trachomatis KDO-related polypeptide;
  - (r) a Chlamydia trachomatis phosphomannomutase-related polypeptide;
  - (s) a Chlamydia trachomatis phosphoglucomutase-related polypeptide;
  - (t) a Chlamydia trachomatis lipid A component-related polypeptide:

- (u) a Chlamydia trachomatis polypeptide that contains an RGD sequence;
- (v) a Chlamydia trachomatis Type III secreted polypeptide;
- (w) a Chlamydia trachomatis cell wall anchored surface polypeptide; or
- (x) a Chlamydia trachomatis polypeptide that is not found in Chlamydia trachomatis.
- 22. A fusion protein encoded by the polynucleotide of Claim 8.
- 23. The fusion protein of Claim 22 which immunoreacts with seropositive serum of an individual infected with *Chlamydia trachomatis*.
  - 24. An antibody that immunospecifically binds to the polypeptide of Claim 19.
  - 25. An antibody that immunospecifically binds to the fusion protein of Claim 22.
- 26. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
  - (a) contacting the sample with a polynucleotide primer of Claim 1, 2, 3, 4, 5, or 6 in the presence of a polymerase enzyme and nucleotides under conditions which permit primer extension; and
  - (b) detecting the presence of primer extension products in the sample in which the detection of primer extension products indicates the presence of *Chlamydia trachomatis* in the sample.
- 27. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
  - (a) contacting the sample with a polynucleotide probe of Claim 1, 2, 3, 4, 5, or 6 under conditions which permit hybridization of complementary base pairs; and
  - (b) detecting the presence of hybridization complexes in the sample in which the detection of hybridization complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 28. A method for the detection and/or identification of *Chlamydia trachomatis* in a biological sample, comprising:
  - (a) contacting the sample with the antibody of Claim 24 under conditions suitable for the formation of immune complexes; and

- (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 29. A method for the detection and/or identification of antibodies to *Chlamydia* trachomatis in a biological sample, comprising:
  - (a) contacting the sample with a polypeptide of Claim 19 under conditions suitable for the formation of immune complexes; and
  - (b) detecting the presence of immune complexes in the sample, in which the detection of immune complexes indicates the presence of *Chlamydia trachomatis* in the sample.
- 30. A DNA chip containing an array of polynucleotides comprising at least one of the polynucleotides of Claim 1, 2, 3, 4, 5, or 6.
- 31. A protein chip containing an array of polypeptides comprising at least one of the polypeptides of Claim 19.
- 32. An immunogenic composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 33. An immunogeneic composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.
- 34. An immunogenic composition comprising the fusion protein of Claim 22 and a pharmaceutically acceptable carrier.
- 35. An immunogenic composition comprising the fusion protein of Claim 23 and a pharmaceutically acceptable carrier.
- 36. A pharmaceutical composition comprising the polypeptide of Claim 19 and a pharmaceutically acceptable carrier.
- 37. A pharmaceutical composition comprising the polypeptide of Claim 20 and a pharmaceutically acceptable carrier.

- 38. A pharmaceutical composition comprising the polypeptide of Claim 22 and a pharmaceutically acceptable carrier.
- 39. A pharmaceutical composition comprising the polypeptide of Claim 23 and a pharmaceutically acceptable carrier.
- 40. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 32.
- 41. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 33.
- 42. A method of immunizing against *Chlamydia trachomatis*, comprising administering to a host an immunizing amount of the immunogenic composition of Claim 34.
- 43. A method of immunizing against *Chlamydia trachomatis*, comprising: administering to a host an immunizing amount of the immunogenic composition of Claim 35.
- 44. A DNA immunogenic composition comprising the expression vector of Claim 11.
- 45. The DNA composition of Claim 44, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia trachomatis*.
- 46. A DNA immunogenic composition comprising the expression vector of Claim 12.
- 47. The DNA composition of Claim 46, wherein the DNA composition directs the expression of a neutralizing epitope of *Chlamydia trachomatis*.
  - 48. A screening assay, comprising:
  - (a) contacting a test compound with an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6; and
  - (b) detecting whether binding occurs.
  - 49. A screening assay, comprising:
  - (a) contacting a test compound with the polypeptide of Claim 19; and

- (b) detecting whether binding occurs.
- 50. A screening assay, comprising:
- (a) contacting a test compound with the polypeptide of Claim 22; and
- (b) detecting whether binding occurs.
- 51. A kit comprising a container containing an isolated polynucleotide of Claim 1, 2, 3, 4, 5 or 6.
  - 52. The kit of Claim 51 wherein the polynucleotide is a primer or a probe.
- 53. The kit of Claim 51 wherein the polynucleotide is a primer and the kit further comprises a container containing a polymerase.
- 54. The kit of Claim 51 which further comprises a container containing deoxynucleotide triphosphates.
- 55. A kit comprising a container containing an antibody that immunospecifically binds to the polypeptide of Claim 19.
- 56. A kit comprising a container containing an antibody that immunospecifically binds to the fusion protein of Claim 22.



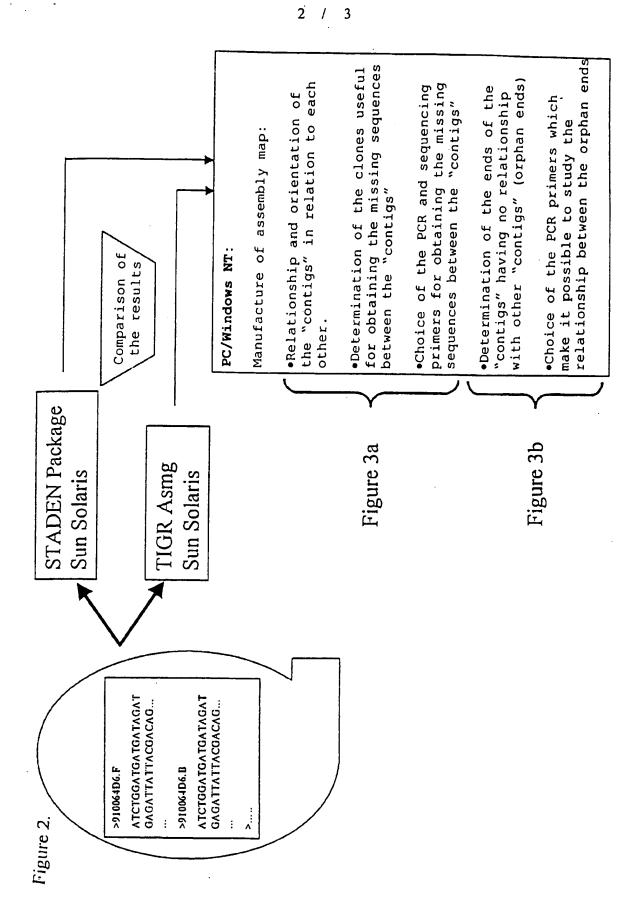


FIGURE 3A

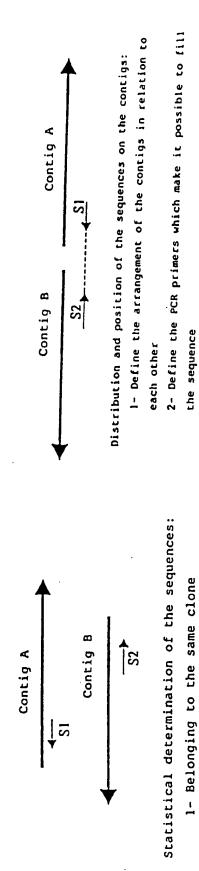
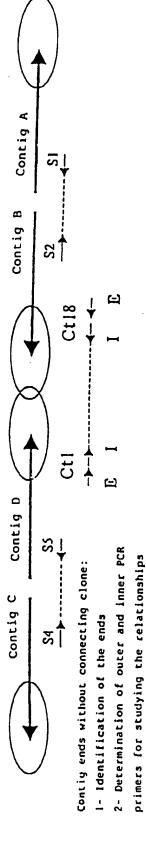


FIGURE 3B



between the contigs E: outer primers

I: inner primers

Situated on two different contigs



#### **PCT**

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#### INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

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(74) Agents: MARTIN, Jean-Jacques et al.; Cabinet Regimbeau, 26, avenue Kléber, F-75116 Paris (FR). (81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

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(54) Title: CHLAMYDIA TRACHOMATIS GENOMIC SEQUENCE AND POLYPEPTIDES, FRAGMENTS THEREOF AND USES THEREOF, IN PARTICULAR FOR THE DIAGNOSIS, PREVENTION AND TREATMENT OF INFECTION

#### (57) Abstract

The subject of the invention is the genomic sequence and the nucleotide sequences encoding polypeptides of Chlamydia trachomatis, such as cellular envelope polypeptides, which are secreted or specific, or which are involved in metabolism, in the replication process or in virulence, polypeptides encoded by such sequences, as well as vectors including the said sequences and cells or animals transformed with these vectors. The invention also relates to transcriptional gene products of the Chlamydia trachomatis genome, such as, for example, antisense and ribozyme molecules, which can be used to control growth of the microorganism. The invention also relates to methods of detecting these nucleic acids or polypeptides and kits for diagnosing Chlamydia trachomatis infection. The invention also relates to a method of selecting compounds capable of modulating bacterial infection and a method for the biosynthesis or biodegradation of molecules of interest using the said nucleotide sequences or the said polypeptides. The invention finally comprises, pharmaceutical, in particular vaccine, compositions for the prevention and/or treatment of bacterial, in particular Chlamydia trachomatis, infections.

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Inter 'onal Application No PCT/IB 98/01939

A. CLASSIF IPC 6	C12N15/31 A01K67/027	MATTER C12N15/62 A61K39/118	C07K14/29 G01N33/53	5 C07K10 C12Q1,		C07K19/00
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C DOCUME	ENTS CONSIDERED TO	BE RELEVANT				
Category *		ith indication, where app	ropriate, of the releva	nt passages		Relevant to claim No.
Y	primary st trachomati J BACTERIO XP00207680 UNITED STA page 5153,		Chlamydia tein." 169 (11) P5	152-6,		1-3,7,9, 11,13, 26,27, 30,44, 45,48, 51-54
Y	-page 5156  TOMB J F E SEQUENCE O HELICOBACT NATURE, vol. 388,	F THE GASTRIC ER PYLORI" 7 August 1997 547, XP002062	olumn, para OMPLETE GEN PATHOGEN (1997-08-0	graph 1 OME		1-3,7,9, 11,13, 26,27, 30,44, 45,48, 51-54
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Intr tional Application No PCT/IB 98/01939

Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
CLARKE IN ET AL: "Molecular cloning and sequence analysis of a developmentally regulated cysteine-rich outer membrane protein from Chlamydia trachomatis." GENE, NOV 30 1988, 71 (2) P307-14, XP002076807 NETHERLANDS abstract	1
GU L ET AL: "Cloning and characterization of a secY homolog from Chlamydia trachomatis."  MOL GEN GENET, MAY 25 1994, 243 (4)  P482-7, XP002076808  GERMANY  abstract  page 486; figures 2,3	1
KAUL R ET AL: "The chlamydial EUO gene encodes a histone H1-specific protease." J BACTERIOL, SEP 1997, 179 (18) P5928-34, XP002076809 UNITED STATES abstract	1
PEDERSEN LB ET AL: "Purification of recombinant Chlamydia trachomatis histone H1-like protein Hc2, and comparative functional analysis of Hc2 and Hc1." MOL MICROBIOL, APR 1996, 20 (2) P295-311, XP002076810 ENGLAND abstract page 295, right-hand column, paragraph 2 -page 307, left-hand column, paragraph 2	1
ZHANG Y ET AL: "Elongation factor Ts of Chlamydia trachomatis: structure of the gene and properties of the protein." ARCH BIOCHEM BIOPHYS, AUG 1 1997, 344 (1) P43-52, XP002076811 UNITED STATES abstract page 43, right-hand column, paragraph 1	1
EP 0 293 079 A (ALBERTA LTD 368800) 30 November 1988 (1988-11-30) claims 1-19	1
	sequence analysis of a developmentally regulated cysteine-rich outer membrane protein from Chlamydia trachomatis."  GENE, NOV 30 1988, 71 (2) P307-14, XP002076807 NETHERLANDS abstract  GU L ET AL: "Cloning and characterization of a secY homolog from Chlamydia trachomatis."  MOL GEN GENET, MAY 25 1994, 243 (4) P482-7, XP002076808 GERMANY abstract page 486; figures 2,3  KAUL R ET AL: "The chlamydial EUO gene encodes a histone H1-specific protease." J BACTERIOL, SEP 1997, 179 (18) P5928-34, XP002076809 UNITED STATES abstract  PEDERSEN LB ET AL: "Purification of recombinant Chlamydia trachomatis histone H1-like protein Hc2, and comparative functional analysis of Hc2 and Hc1."  MOL MICROBIOL, APR 1996, 20 (2) P295-311, XP002076810 ENGLAND abstract page 295, right-hand column, paragraph 2 -page 307, left-hand column, paragraph 1 UNITED STATES abstract page 43, right-hand column, paragraph 1  EP 0 293 079 A (ALBERTA LTD 368800) 30 November 1988 (1988-11-30)

Inter onal Application No PCT/IB 98/01939

	ion) DOCUMENTS CONSIDERED TO BE RELEVANT  Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
Category '	Citation of accument, with indication, where appropriate, of the resevant passages			
<b>у,</b> х	STEPHENS RS ET AL: "Genome sequence of an obligate intracellular pathogen of humans: Chlamydia trachomatis [see comments]" SCIENCE, OCT 23 1998, 282 (5389) P754-9, XP002104802 UNITED STATES the whole document	1-3,7,9, 11,13		
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national application No.

PCT/IB 98/01939

Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inter	rnational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:  Remark: Although claims 40-43  are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	rnational Searching Authority found multiple inventions in this international application, as follows:
see	ADDITIONAL SHEET
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. X	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:  1-3 and 7,9,11,13,26,27,30,44,45,48 (partially)
Remark	on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

#### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Invention 1: claims 1-3 and 7,9,11,13,26,27,30,44,45,48 (partially)

nucleotide seq.id.n.l coding for the genome of Chlamydia trachomatis, corresponding

vector , host method of detection , DNA chip , screening

assay and kit .

Invention 2 : claims 4-56 (partially )

ORF2 of Chlamydia trachomatis fragments, corresponding polypeptides, nucleotide

sequences, DNA chip, cloning vector, host, method

for producing polypeptides ,fusion polypeptide , method for the detection , kit , antibody , immunogenic and pharmaceutical composition ,

screening assay .

Inventions 3-1197 : identical to invention 2 , but applied to orf3-1197 , in which invention 3 is limited to ORF3 invention 4 to ORF4 , etc. ... until invention 1197 is limited to ORF1197 .

rmation on patent family members

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